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54 **Alpha-aryl-alpha-phenylethyl-1H-1,2,4-triazole-1-propanenitriles.**

57 This invention relates to certain substituted and unsubstituted alpha-aryl-alpha-phenylethyl-1H-1,2,4-triazole-1-propanenitriles, their enantiomorphs, acid addition salts and metal salt complexes which are active broad-spectrum systemic fungicides effective in controlling phytopathogenic fungi such as corn helminthosporium, barley helminthosporium, cucumber downy mildew, cucumber powdery mildew, cucumber anthracnose, botrytis, tomato early blight, tomato late blight, grape downy mildew, bean powdery mildew, peanut cercospora, rice blast, rice sheath blight, wheat leaf rust, wheat septoria nodorum, wheat powdery mildew and wheat stem rust. The invention also concerns a method of preparing alpha-aryl-alpha-triazolylmethyl-acetonitrile compounds by reacting an aryl acetonitrile compound with a halomethyltriazole.

Description

ALPHA-ARYL-ALPHA-PHENYLETHYL-1H-1,2,4-TRIAZOLE-1-PROPANENITRILES

This invention is concerned with alpha-aryl-alpha-phenyl-ethyl-1H-1,2,4-triazole-1-propanenitriles compositions containing them and their use in controlling phytopathogenic fungi. In the compounds of the invention, the phenethyl and aryl moieties may be substituted or unsubstituted.

Miller et al., U.S. Patent No. 4,363,165 discloses 1- and 4-arylcyanoalkyl-1,2,4-triazoles and their use against phytopathogenic fungi. However, neither the phenethyl triazoles of the present invention nor the benzyl triazoles were made by Miller et al. Therefore the prior art fails to recognize the particular class of compounds to which the present invention is limited or that this class has a surprisingly high degree of fungicidal activity. The phenethyl triazoles of the present invention are not only effective against wheat powdery mildew, wheat stem rust and wheat leaf rust but are significantly superior to the benzyl triazoles and phenyl triazoles in their activity against barley helminthosporium, rice blast and peanut early leafspot.

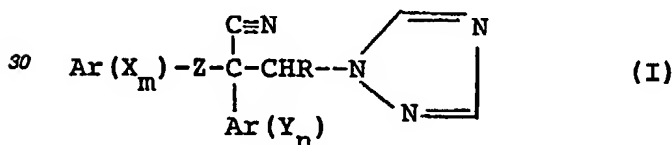
European Patent Application 52,424, published May 26, 1982, generically discloses a class of compounds which embraces the compounds of the present invention. However, none of the present compounds was made or specifically disclosed. In fact, none of the examples of the 52,424 application have a cyano group attached to the quaternary carbon, but include either a hydroxy, methoxy, butoxy or allyloxy group. The three phenethyl triazole compounds which were made and disclosed in the 52,424 application are 4,4-dimethyl-3-hydroxy-3-(1,2,4-triazol-1-yl)methyl-1-(halogen substituted phenyl)-pentanes.

Published German patent application No. 3,216,301 discloses the fungicidal activity of alkoxytriazole-propanitriles.

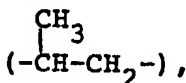
European application 63,099 discloses the use of chloromethyltriazole to produce a 1H-1,2,4-triazol-1-yl-methylphosphonium salt.

Published British patent application 2,119,374 discloses a process to make alpha-(alkoxy, alkenoxy, alkynoxy or phenylalkoxy)-alpha-aryl-triazolylmethyl-acetonitrile compounds.

We have now found a new class of triazole propanenitriles of surprising fungicidal activities. This class comprises alpha-aryl-alpha-phenylethyl-1H-1,2,4-triazole-1-propanenitriles of the formula (I):



wherein Z is an ethylene group ($-\text{CH}_2 - \text{CH}_2 -$), an ethenylene group ($-\text{CH}=\text{CH}-$), an ethynylene group ($-\text{C}\equiv\text{C}-$) or an isopropylene group



which ethylene,

ethenylene or isopropylene groups may optionally be halogenated; $\text{Ar(X}_m\text{)}$ is a substituted or unsubstituted $\text{C}_6 - \text{C}_{10}$ aromatic ring structure; $\text{Ar(Y}_n\text{)}$ is a substituted or unsubstituted aryl; X and Y may be the same or different and may be SO_2Z wherein Z is (C_1 to C_6) alkyl or aryl and a is 0, 1 or 2 halogen, ($\text{C}_1 - \text{C}_6$) alkyl optionally substituted with up to 3 halogens, ($\text{C}_2 - \text{C}_6$) alkenyl optionally substituted with up to 3 halogens, hydroxy, ($\text{C}_1 - \text{C}_6$) alkoxy, (C_2 to C_6) alkenoxy, phenyl optionally substituted with one or 2 halogens, cyano, amino, monoalkylamino having up to six carbon atoms, dialkylamino having up to six carbon atoms in each alkyl group, $-\text{C(O)H}$ or $-\text{C(O)NR}_1\text{R}_2$ wherein R_1 and R_2 are independently hydrogen or alkyl; R is hydrogen or phenyl optionally substituted with up to 3 halogen atoms, a trifluoromethyl or a (C_1 to C_6) alkyl group; and m and n are independently 0 to 3. The new class of triazole propanenitriles also includes the agronomically acceptable enantiomorphs, acid addition salts and metal complexes of the compounds of formula (I).

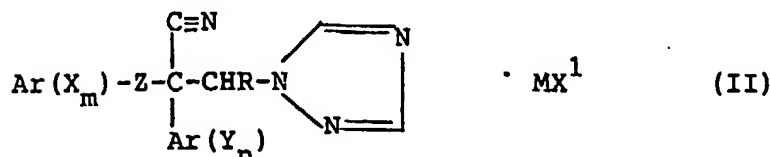
The term "aryl" is meant to include $\text{C}_6 - \text{C}_{10}$ aromatic ring structures, a five member aromatic ring having 4 carbon atoms and one nitrogen, oxygen or sulfur atom, or a six member aromatic ring having 1 or 2 nitrogen atoms and 5 or 4 carbon atoms; all optionally substituted with up to 3 halogen atoms, a trifluoromethyl group or a C_1 to C_6 alkyl group.

The term "alkyl" is meant to include both branched and straight chained alkyl groups of carbon atoms, preferably $\text{C}_1 - \text{C}_8$. Typical alkyl groups which are encompassed by the use of this term include methyl, ethyl, n-propyl, iso-propyl, n-butyl, sec-butyl, iso-butyl, tert-butyl, pentyl, neo-pentyl, iso-pentyl, hexyl, heptyl and iso-octyl.

The term "alkoxy," as used in the present specification, is meant to include alkenoxy, as well as the radical consisting of an alkyl group bonded to an oxygen atom. The preferred groups include (C -C)-alkoxy. Typical alkoxy groups which are encompassed by the use of the term include methoxy, ethoxy, propoxy, n-butoxy, iso-butoxy, pentoxy, hexoxy and allyloxy.

The acids which can be utilized in making the acid addition salts of the present invention include, for example, hydrochloric, hydrobromic, nitric, sulfuric, phosphoric, hydriodic, hydrofluoric, perchloric, p-toluenesulfonic, methanesulfonic, acetic, citric, tartaric, malic, maleic, oxalic, fumaric and phthalic acids.

Included within this invention are the metal salt complexes of the formula (II):



wherein Z, Ar(Y_n), X, Y, n and m are as defined in formula (I) above and M is a cation selected from Group IIA, IB, IIB, VIB, VIIIB and VIII of the Periodic Table and X¹ is an anion selected so that the sum of the valence charges of the cation M and anion X¹ equal zero.

Typical cations encompassed by this invention are magnesium, manganese, copper, nickel, zinc, iron, cobalt, calcium, tin, cadmium, mercury, chromium, lead and barium.

Typical anions encompassed by this invention are chloride, bromide, iodide, fluoride, sulfate, bisulfate, perchlorate, nitrate, nitrite, phosphate, carbonate, bicarbonate, acetate, citrate, oxalate, tartrate, malate, maleate, fumarate, p-toluenesulfonate, methanesulfonate, mono- or di-(C₁ -C₄) alkylidithiocarbamate and (C₁ -C₄) alkylenebisdithiocarbamate.

In one preferred aspect this invention comprises the compounds, agronomically acceptable enantiomorphs, salts and complexes of formulas (I) and (II) wherein Z is an ethylene group, Ar is phenyl and X and Y are hydrogen, halogen or trifluoromethyl. In another more preferred embodiment Y_n is hydrogen, 2-alkoxy, 4-halo or 3-trifluoromethyl and X is a hydrogen, 4-halo or 4-trifluoromethyl. In yet another preferred embodiment, Y is 2-halo and X is 4-halo or Y is 4-halo and X is 2-halo. In yet another preferred embodiment, X is 3-halo and Y is hydrogen.

In another preferred embodiment Z is ethynylene, Ar is phenyl and X and Y are hydrogen, halogen or trifluoromethyl. In a more preferred embodiment Z is ethynylene and Ar(X_m) is a halogen substituted phenyl.

Typical compounds encompassed by the present invention which were made include the following:

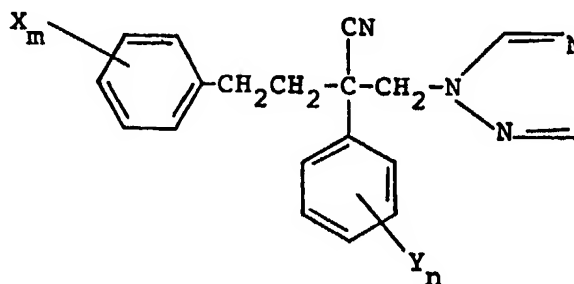
1. alpha-(4-chlorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
2. alpha-phenyl-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
3. alpha-(2-methoxyphenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
4. alpha-(4-fluorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
5. alpha-(2,4-dichlorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
6. alpha-(4-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
7. alpha-(4-chlorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
8. alpha-(4-chlorophenyl)-alpha-[2-(4-methylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
9. alpha-(4-chlorophenyl)-alpha-[2-(4-methoxyphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
10. alpha-[2-(4-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
11. alpha-[2-(4-fluorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
12. alpha-(2-phenylethyl)-alpha-(4-phenylphenyl)-1H-1,2,4-triazole-1-propanenitrile
13. alpha-phenyl-alpha-[2-(2-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
14. alpha-phenyl-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
15. alpha-[2-(2,4-dichlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
16. alpha-[2-(4-bromophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
17. alpha-[2-(2-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
18. alpha-[2-(3-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
19. alpha-phenyl-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
20. alpha-(4-chlorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
21. alpha-(4-fluorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
22. alpha-[2-(4-bromophenyl)ethyl]-alpha-(4-chlorophenyl)-1H-1,2,4-triazole-1-propanenitrile
23. alpha-(4-chlorophenyl)-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
24. alpha-(4-fluorophenyl)-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
25. alpha-[2-(4-bromophenyl)ethyl]-alpha-(4-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
26. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(4-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
27. alpha-[2-(3-bromophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
28. alpha-(4-fluorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
29. alpha-(2-methoxyphenyl)-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
30. alpha-[2-(3-chlorophenyl)ethyl]-alpha-(4-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
31. alpha-(4-chlorophenyl)-alpha-[2-(3-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile

32. alpha-(4-bromophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
33. alpha-(4-bromophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
34. alpha-(3-chlorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
35. alpha-(3-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
36. alpha-(3-chlorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
37. alpha-(4-bromophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
38. alpha-[2-(3-bromophenyl)ethyl]-alpha-(4-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
39. alpha-[2-(3-bromophenyl)ethyl]-alpha-(4-chlorophenyl)-1H-1,2,4-triazole-1-propanenitrile
40. alpha-(2-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
41. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
42. alpha-(4-chlorophenyl)-alpha-[2-(2-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
43. alpha-(4-fluorophenyl)-alpha-[2-(2-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
44. alpha-(2-chlorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
45. alpha-(2-fluorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
46. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(3-trifluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
47. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(3-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
48. alpha-(2-bromophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
49. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
50. alpha-(2-phenylethyl)-alpha-(3-trifluoromethylphenyl)-1H-1,2,4-triazole-1-propanenitrile
51. alpha-(3-trifluoromethylphenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
52. alpha-(3-fluorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
53. alpha-(3-chlorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
54. alpha-(2-bromophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
55. alpha-(2-bromophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
56. alpha-(3-fluorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
57. alpha-(2-chlorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
58. alpha-[2-(2-methoxyphenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
59. alpha-[2-(3-methoxyphenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
60. alpha-[2-(3,4-dimethoxyphenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
61. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(4-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
62. alpha-(4-methoxyphenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
63. alpha-(2-chloro-6-fluorophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
64. alpha-(2-chloro-6-fluorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
65. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2,6-dichlorophenyl)-1H-1,2,4-triazole-1-propanenitrile
66. alpha-[2-(3-bromophenyl)ethyl]-alpha-(3-chlorophenyl)-1H-1,2,4-triazole-1-propanenitrile
67. alpha-[2-(4-bromophenyl)ethyl]-alpha-(3-chlorophenyl)-1H-1,2,4-triazole-1-propanenitrile
68. alpha-[2-(3-bromophenyl)ethyl]-alpha-(3-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
69. alpha-[2-(4-bromophenyl)ethyl]-alpha-(3-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
70. alpha-phenyl-alpha-(2-phenyl)propyl-1H-1,2,4-triazole-1-propanenitrile
71. alpha-(3-fluorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
72. alpha-[2-(4-fluorophenyl)ethyl]-alpha-(3-trifluoromethylphenyl)-1H-1,2,4-triazole-1-propanenitrile
73. alpha-[2-(1-naphthyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
74. alpha-[2-(4-bromophenyl)ethyl]-alpha-(2-ethoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
75. alpha-[2-(4-bromophenyl)ethyl]-alpha-(3-trifluoromethylphenyl)-1H-1,2,4-triazole-1-propanenitrile
76. alpha-(2-ethoxyphenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
77. alpha-(2-ethoxyphenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
78. alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2-ethoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
79. alpha-(2-ethoxyphenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
80. alpha-[2-(3,4-dichlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
81. alpha-(3-chlorophenyl)-alpha-[2-(3-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
82. alpha-(3-chlorophenyl)-alpha-[2-(3-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
83. alpha-[2-(3-fluorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
84. alpha-[2-(3-chlorophenyl)ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
85. alpha-(4-bromophenyl)-alpha-[2-(3-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
86. alpha-(4-chlorophenyl)-alpha-[2-(3-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
87. alpha-(3-bromophenyl)-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile
88. alpha-[2-(3-bromophenyl)ethyl]-alpha-(2-ethoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile
89. alpha-(3-fluorophenyl)-alpha-[2-(3-fluorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
90. alpha-(4-bromophenyl)-alpha-[2-(3-bromophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile
91. alpha-[2-(3-chlorophenyl)ethyl]-alpha-(3-fluorophenyl)-1H-1,2,4-triazole-1-propanenitrile
92. alpha-[2-(3,5-dichlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
93. alpha-[2-(4-methoxyphenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile
94. alpha-[2-(4-chlorophenyl)ethyl]-alpha-phenyl-beta-phenyl-1H-1,2,4-triazole-1-propanenitrile (one isomer at beta position)

95. α -[2-(4-chlorophenyl)ethyl]- α -(2-thienyl)-1H-1,2,4-triazole-1-propanenitrile
 96. α -[2-(4-chlorophenyl)ethyl]- α -(2-pyridyl)-1H-1,2,4-triazole-1-propanenitrile
 97. α -phenyl- α -(e)(2-phenylethenyl)-1H-1,2,4-triazole-1-propanenitrile
 98. α -phenyl- α -(e)[2-(4-chlorophenyl)ethenyl]-1H-1,2,4-triazole-1-propanenitrile

The structures of the Compounds 1-96 are set forth in Table 1 below.

TABLE 1

CompoundX_mY_n

1

H

4Cl

2

H

H

3

H

2OCH₃

0251775

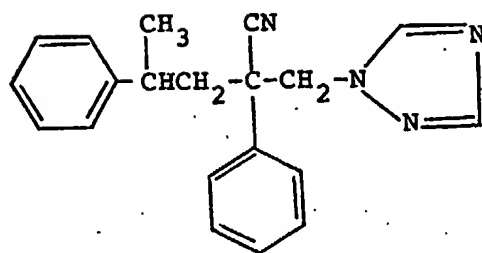
<u>Compound</u>	<u>X_n</u>	<u>Y_n</u>
4	H	4F
5	H	2,4Cl
6	4Cl	4Cl
7	4F	4Cl
8	4CH ₃	4Cl
9	4OCH ₃	4Cl
10	4Cl	H
11	4F	H
12	H	4Ø
13	2CF ₃	H
14	3CF ₃	H
15	2,4Cl	H
16	4Br	H
17	2Cl	H
18	3Cl	H
19	4CF ₃	H
20	3CF ₃	4Cl
21	3CF ₃	4F
22	4Br	4Cl
23	4CF ₃	4Cl
24	4CF ₃	4F
25	4Br	4F
26	4Cl	4F
27	3Br	H
28	4F	4F
29	4CF ₃	2OCH ₃
30	3Cl	4F
31	3Cl	4Cl
32	H	4Br
33	4F	4Br
34	H	3Cl
35	4Cl	3Cl
36	4F	3Cl
37	4Cl	4Br

0 251 775

<u>Compound</u>	<u>X_m</u>	<u>Y_n</u>
38	3Br	4F
39	3Br	4Cl
40	4Cl	2Cl
41	4Cl	2F
42	2F	4Cl
43	2F	4F
44	4F	2Cl
45	4F	2F
46	4Cl	3CF ₃
47	4Cl	3F
48	4Cl	2Br
49	4Cl	2OCH ₃
50	H	3CF ₃
51	3CF ₃	3CF ₃
52	H	3F
53	3CF ₃	3Cl
54	H	2Br
55	3CF ₃	2Br
56	3CF ₃	3F
57	3CF ₃	2Cl
58	2OCH ₃	H
59	3OCH ₃	H
60	3,4OCH ₃	H
61	4Cl	4OCH ₃
62	H	4OCH ₃
63	H	2Cl,6F
64	4Cl	2Cl,6F
65	4Cl	2,6Cl
66	3Br	3Cl
67	4Br	3Cl
68	3Br	3F
69	4Br	3F
70	See formula (III) below	
71	4F	3F

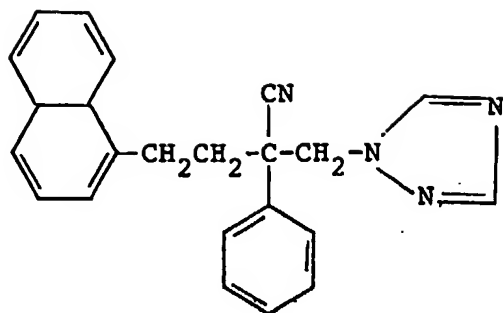
<u>Compound</u>	<u>X_m</u>	<u>Y_n</u>
72	4F	3CF ₃
73	See formula (IV) below	
74	4Br	20Et
75	4Br	3CF ₃
76	3CF ₃	20Et
77	H	20Et
78	4Cl	20Et
79	4F	20Et
80	3,4Cl	H
81	3Cl	3Cl
82	3F	3Cl
83	3F	H
84	3Cl	2OCH ₃
85	3Cl	4Br
86	3F	4Cl
87	H	3Br
88	3Br	20Et
89	3F	3F
90	3Br	4Br
91	3Cl	3F
92	3,5Cl	H
93	4OCH ₃	H
94	See formula (V) below	
95	See formula (VI) below	
96	See formula (VII) below	

70.



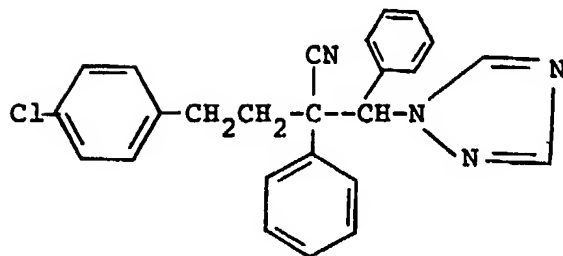
(III)

73.



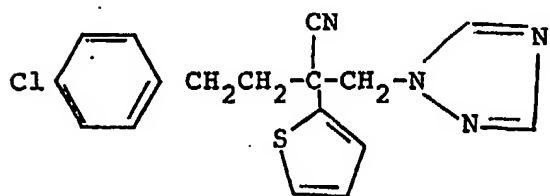
(IV)

94.



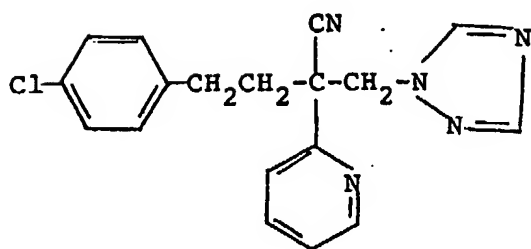
(V)

95.



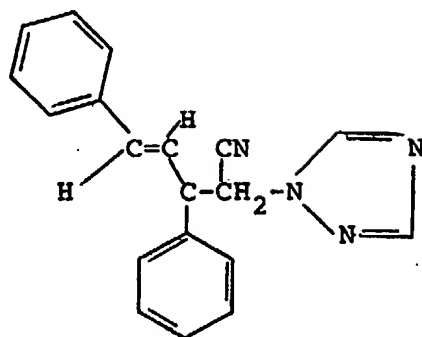
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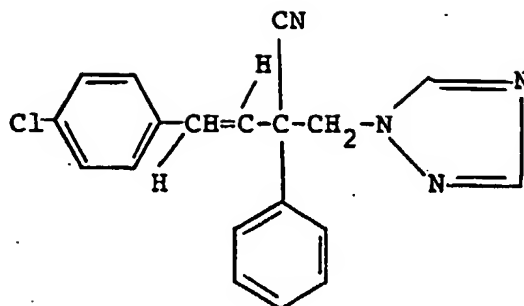


(VII)

97.



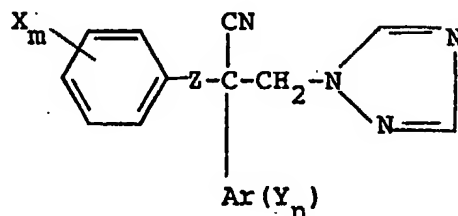
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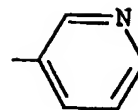
(Compounds 97 and 98 are the E-isomers)

Other examples which are included in the present invention are set forth in Table 2.

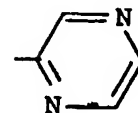
TABLE 2



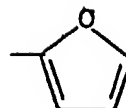
Compound	Z	X _m	Ar (Y _n)
99	-CH ₂ CH ₂ -	-H	-Ø (4CH ₂ CH ₂ CH ₂ Cl)
100	-CH ₂ CH ₂ -	4CH=CHF	-Ø
101	-CH ₂ CH ₂ -	-Ø (2,4Cl)	-Ø
102	-CH ₂ CH ₂ -	4CN	-Ø (3CN)
103	-CH ₂ CH ₂ -	4C(O)H	-Ø
104	-CH ₂ CH ₂ -	4NHCH ₃	-Ø
105	-CH ₂ CH ₂ -	2SCH ₃	-Ø (4SO ₂ CH ₃)
106	-CH=CH-	-H	-Ø (3Cl)
107	-C=C-	-H	-Ø (4Cl)
108	-CH ₂ CH ₂ -	4S(O)CH ₂ CH ₃	-Ø
109	-CHClCH ₂ -	4Cl	-Ø
110	-CHClCHCl-	4F	-Ø
111	-CHBrCH ₂ -	4Br	-Ø

112 $-\text{CH}_2\text{CH}_2-$ 4Cl

5

113 $-\text{CH}_2\text{CH}_2-$ 4F

10

114 $-\text{CH}_2\text{CH}_2-$ 4Br

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Comparative compounds which were made and tested include:

C2a. alpha,alpha-diphenyl-1H-1,2,4-triazole-1-propanenitrile

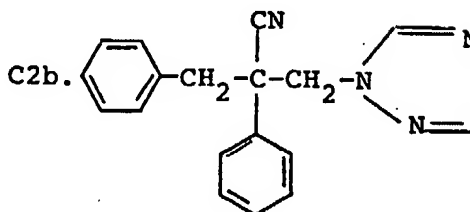
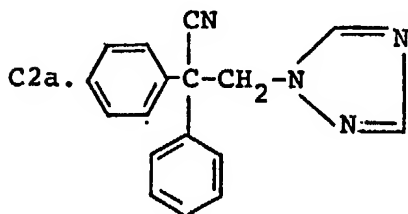
C2b. alpha-benzyl-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile

C6. alpha-(4-chlorobenzyl)-alpha-(4-chlorophenyl)-1H-1,2,4-triazole-1-propanenitrile

C10. alpha-(4-chlorobenzyl)-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile

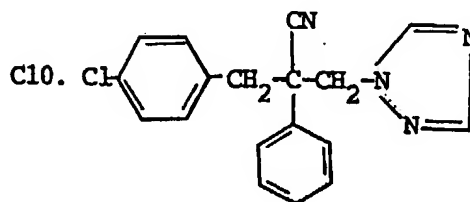
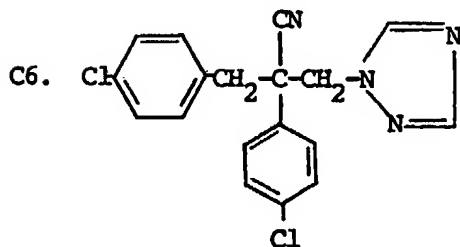
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The structures of the Comparative compounds are set forth below.



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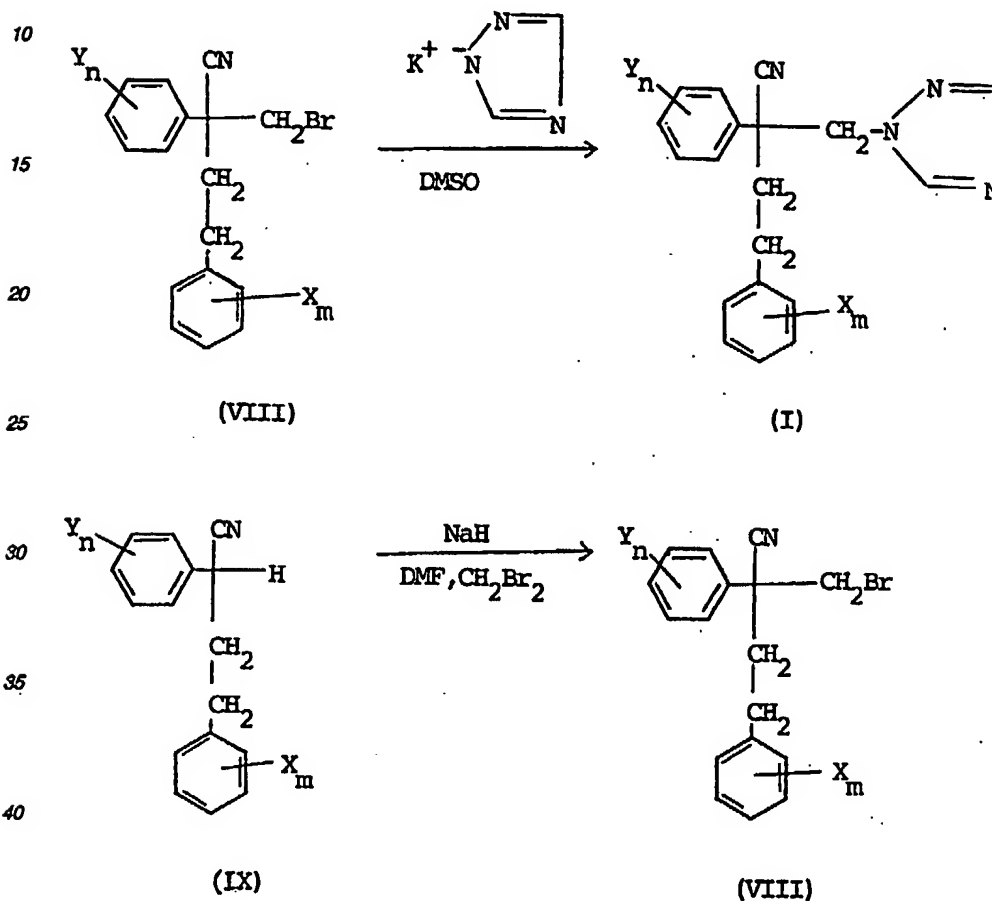
The triazoles of the present invention can be prepared by conventional synthesis routes. For example, the triazoles may be prepared by nucleophilic displacement of the alpha-(bromomethyl)-alpha-(2-phenylethyl)-phenylacetonitrile (VIII) by a salt, preferably an alkali metal salt, of the triazole, generally about 1 to about 3 equivalents. This reaction can be run either neat or, preferably, in an appropriate solvent such as dimethylsulfoxide (DMSO), N,N-dimethylformamide (DMF), toluene or xylene at a temperature from about 0°C to about 150°C, preferably from about 25°C to about 100°C. The compound VIII is prepared by bromomethylation of the alpha-2-phenylethyl-phenylacetonitrile (IX) by methylenabromide (generally about 1.1 to about 2 equivalents) under basic conditions, e.g., sodium or potassium hydroxide, sodium or potassium hydride, potassium methoxide and potassium-t-butoxide (generally about 1.1 to about 2 equivalents) preferably with the use of a solvent such as DMSO with sodium hydroxide or DMF with the hydrides and oxides, at a temperature from about 0°C to about 150°C, preferably from about 25°C to about 100°C. The compound IX can be prepared by phase transfer alkylation of the appropriately substituted benzylcyanides (X) with generally about 1 to about 2 equivalents of a 2-phenylethylmethanesulfonate (mesylate) or

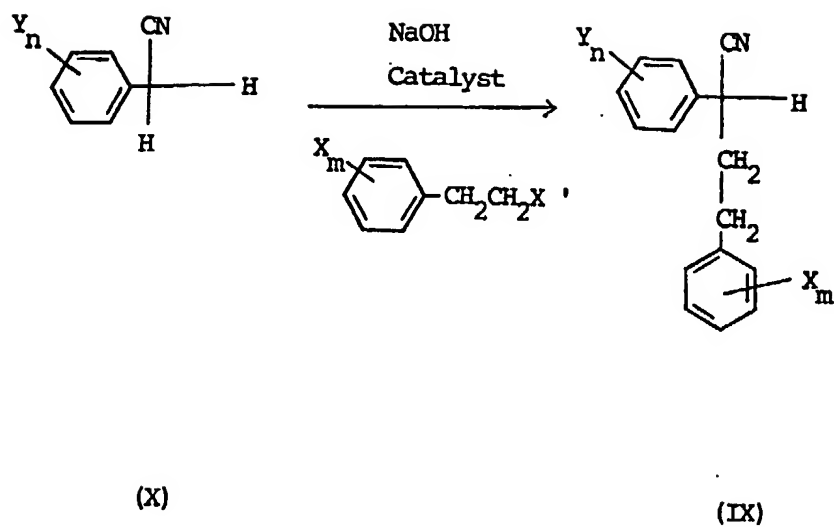
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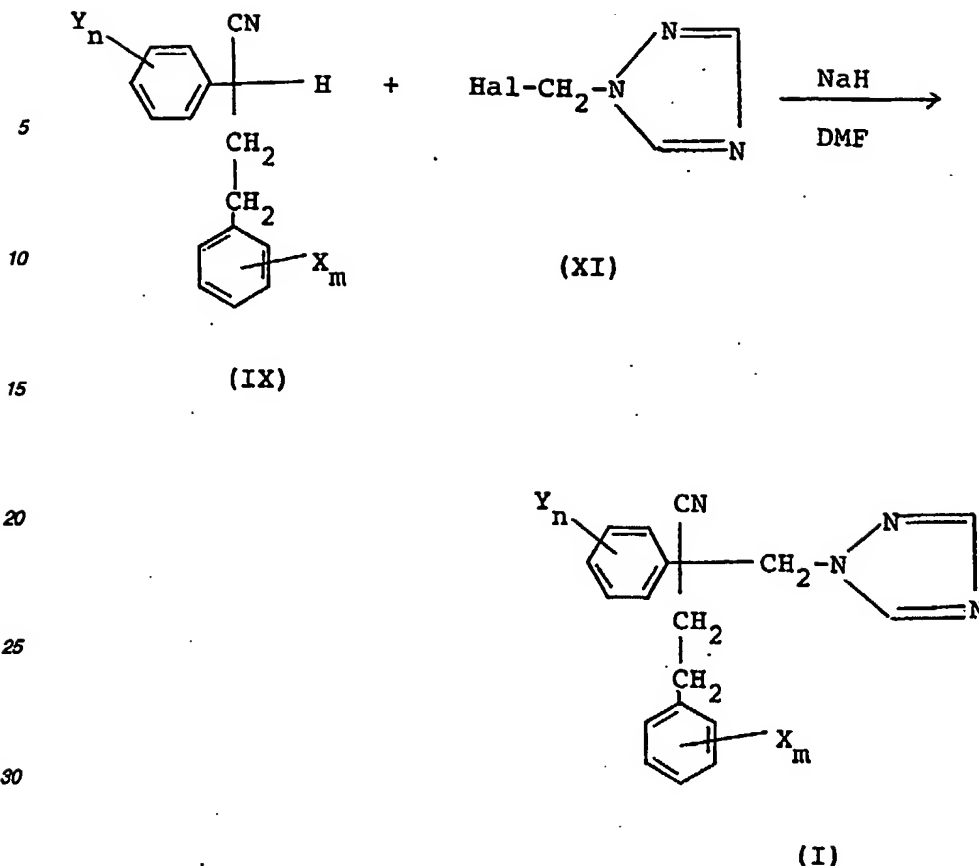
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p-toluenesulfonate (tosylate) in the presence of a strong base, e.g., 50% (w/w) sodium hydroxide, and a catalyst, e.g., tetrabutylammonium bromide (TBABr). Compound IX can also be prepared by alkylation of the appropriately substituted benzylcyanide with about 1 to 2 equivalents of a 2-phenylethylhalide in the presence of a strong base such as metal hydrides, e.g. sodium or potassium hydride, using DMF or DMF/toluene as the solvent. Both the benzylcyanides and the alkyl halides can be readily prepared by techniques known from the literature. This synthesis scheme is shown below:





The phenethyl triazole (I) can be prepared directly from the phenylethyl-phenylacetonitrile (IX) in one step by the reaction with a halomethyltriazole (XI) in a solvent such as DMF. Two equivalents of a strong base such as sodium hydride or potassium hydride are used. This synthesis scheme is shown below:



This reaction is not limited to the preparation of the triazoles of the present invention. The reaction can also be used to prepare compounds having a quaternary carbon bonded to an aryl group, a cyano group and a triazole group such as disclosed in U.S. patent no. 4,366,165 and British patent application 2,119,374. The method disclosed in the prior art requires the additional step of preparing an aryloxy-(halomethyl, alkylsulphonyloxymethyl or arylsulphonyloxymethyl) compound which is reacted with a triazole or an alkali metal triazole derivative. This extra step is eliminated by the process of the present invention.

The acid addition salts of the triazoles of this invention can be prepared by standard techniques wellknown in the art. For example, the triazole of formula (I) can be dissolved in an appropriate solvent such as diethyl ether, tetrahydrofuran, ethanol, methanol and the like or combinations thereof and treated with an equivalent or excess amount of a mineral or organic acid which may or may not be dissolved in an appropriate solvent, the mixture is then either cooled or evaporated to give the salt which can either be used as such or recrystallized from an appropriate solvent or combination of appropriate solvents.

The metal salt complexes of the above triazoles of this invention can be prepared by adding dropwise, with stirring, a stoichiometric amount of a metal salt dissolved in an appropriate solvent or combination of solvents to a solution of the triazole of formula (I) dissolved in a similarly appropriate solvent or combination of solvents. The reaction mixture is briefly stirred and the solvent is removed under reduced pressure to give the metal salt complex of the respective triazoles of formula (II).

The metal salt complexes can also be prepared by mixing stoichiometric or excess amounts of the metal salt and a triazole of formula (I) in the desired amount of solvent containing the appropriate adjuvants just prior to spraying the plants. Adjuvants that may be included in this "in situ" preparation may be detergents, emulsifiers, wetting agents, spreading agents, dispersing agents, stickers, adhesives, and the like which are used in agricultural applications.

Solvents that can be utilized in these procedures include any polar solvent, e.g., water, methanol, ethanol, isopropanol or ethylene glycol and any aprotic dipolar solvent, e.g., dimethylsulfoxide, acetonitrile, dimethylformamide, nitromethane or acetone.

The metal salt cations that can be used in these procedures can be selected from the group consisting of calcium, magnesium, manganese, copper, nickel, zinc, iron, cobalt, tin, cadmium, mercury, chromium, lead, barium and the like.

Any appropriate anion, e.g., chloride, bromide, iodide, sulfate, bisulfate, phosphate, nitrate, perchlorate, carbonate, bicarbonate, hydrosulfide, hydroxide, acetate, oxalate, malate, citrate and the like may be utilized

as the counterion in the metal salt.

The compounds of this invention possess an asymmetric carbon atom and thus exist as racemic mixtures. The d and l enantiomorphs in these racemic mixtures can be separated via standard techniques such as fractional crystallization with d-tartaric acid, l-tartaric acid, l-quinic acid and the like followed by basification and extraction of the d or l enantiomorph free base.

The enantiomorphs, acid addition salts and metal salt complexes of the present invention are useful as agricultural fungicides and as such can be applied to various loci such as the seed, the soil or the foliage. For such purposes these compounds can be used in the technical or pure form as prepared, as solutions or as formulations. The compounds are usually taken up in a carrier or are formulated so as to render them suitable for subsequent dissemination as fungicides. For example, these chemical agents can be formulated as wettable powders, emulsifiable concentrates, dusts, granular formulations, aerosols, or flowable emulsion concentrates. In such formulations, the compounds are extended with a liquid or solid carrier and, when desired, suitable surfactants are incorporated.

It is usually desirable, particularly in the case of foliar spray formulations, to include adjuvants, such as wetting agents, spreading agents, dispersing agents, stickers, adhesives and the like in accordance with agricultural practices. Such adjuvants commonly used in the art can be found in the John W. McCutcheon, Inc. publication "Detergents and Emulsifiers, Annual".

In general, the compounds of this invention can be dissolved in certain solvents such as acetone, methanol, ethanol, dimethylformamide or dimethyl sulfoxide and such solutions can be extended with water. The concentrations of the solution can vary from about 1% to about 90% with a preferred range being from about 5% to about 50%.

For the preparation of emulsifiable concentrates, the compound can be dissolved in suitable organic solvents, or a mixture of solvents, together with an emulsifying agent which permits dispersion of the fungicide in water. The concentration of the active ingredient in emulsifiable concentrates is usually from about 10% to about 90% and in flowable emulsion concentrates, this can be as high as about 75%.

Wettable powders suitable for spraying, can be prepared by admixing the compound with a dispersing agent and a finely divided solid, such as clays, inorganic silicates and carbonates, and silicas, and optionally incorporating wetting agents and sticking agents in such mixtures. The concentration of active ingredients in such formulations is usually in the range of from about 20% to about 98%, preferably from about 40% to about 75%. A typical wettable powder is made by blending 50 parts of alpha-[2-(4-chlorophenyl) ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile, 45 parts of a synthetic precipitated hydrated silicon dioxide sold under the trademark Hi-Sil, 1 part sodium lauryl sulfate and 5 parts of sodium lignosulfonate. In another preparation a kaolin type (Barden) clay is used in place of the Hi-Sil in the above wettable powder, and in another such preparation 25% of the Hi-Sil is replaced with a synthetic sodium silico aluminate sold under the trademark Zeolox 7.

Dusts are prepared by mixing the triazoles, enantiomorphs, salts and complexes thereof with finely divided inert solids which can be organic or inorganic in nature. Materials useful for this purpose include botanical flours, silicas, silicates, carbonates and clays. One convenient method of preparing a dust is to dilute a wettable powder with a finely divided carrier. Dust concentrates containing from about 20% to about 80% of the active ingredient are commonly made and are subsequently diluted to from about 1% to about 10% use concentration.

The enantiomorphs, salts and complexes thereof can be applied as fungicidal sprays by methods commonly employed, such as conventional high-gallonage hydraulic sprays, low-gallonage sprays, air-blast spray, aerial sprays and dusts. The dilution and rate of application can be readily determined by one in the art depending upon the type of equipment used, the desired method, timing and frequency of applications, plants to be treated and diseases to be controlled. Generally, however, the fungicidal compounds of the present invention will be applied in an amount of from about 1.12 kg to about 2242 kg (0.01 to 20 pounds) of active ingredient per square kilometer (per acre) when applied foliarly or to the soil.

As a seed protectant, the amount of the compound coated on the seeds is usually from about 3.13 to about 250 grams (0.05 to 4 ounces) of active ingredient per hundred kilograms (per hundred pounds) of seed and preferably from 6.25 to about 62.5 grams (0.1 to 1 ounce) per hundred kilograms (per hundred pounds) of seed. As a soil fungicide the compounds can be incorporated in the soil or applied to its surface usually at a rate of from about 5.6 to about 2242 kg (0.05 to 20 pounds), preferably from about 2.24 kg to about 1121 kg (0.02 to 10 pounds) and more preferably from about 11.21 kg to about 336.3 kg (0.1 to 3 pounds) of active ingredient square kilometer (per acre). As a foliar fungicide, the compounds are usually applied to growing plants at a rate of from about 1.12 kg to about 1121 kg (0.01 to 10 pounds), preferably from about 2.24 kg to about 560.5 kg (0.02 to 5) and more preferably from about 3.36 kg to about 112.1 kg (0.03 to 1 pound) of active ingredient square kilometer (per acre).

Fungicides which can be combined with the fungicides of this invention include:

(a) dithiocarbamate and derivatives such as: ferric dimethyldithiocarbamate (ferbam), zinc dimethyldithiocarbamate (ziram), manganese ethylenebisdithiocarbamate (maneb) and its coordination product with zinc ion (mancozeb), zinc ethylenebisdithiocarbamate (zineb), zinc propylenebisdithiocarbamate (propineb), sodium methyldithiocarbamate (metham), tetramethylthiuram disulfide (thiram), the complex of zineb and polyethylene thiuram disulfide, 3,5-dimethyl-1,3,5-2H-tetrahydrothiadiazine-2-thione (dazomet); and mixtures of these and mixtures with copper salts;

(b) nitrophenol derivatives such as:

dinitro-(1-methylheptyl) phenyl crotonate (dinocap), 2-sec-butyl-4,6-dinitrophenyl 3,3-dimethylacrylate (binapacryl), and 2-sec-butyl-4,6-dinitrophenyl isopropyl carbonate;

(c) heterocyclic structures such as: Systhane (a trademark of Rohm and Haas for myclobutanil), tridemifon, N-trichloromethylthiotetrahydrophthalimide (captan), N-trichloromethylthiophthalimide (folpet), 2-heptadecyl-2-imidazole acetate (glydine), 2-octylisothiazol-3-one, 2,4-dichloro-6-(o-chloroanilino)-s-triazine, diethyl phthalimidophosphorothioate, 4-butyl-1,2,4-triazole, 5-amino-1[bis(dimethylamino)-phosphinyl]-3-phenyl-1,2,4-triazole, 5-ethoxy-3-trichloromethyl-1,2,4-thiadiazole, 2,3-dicyano-1,4-dithiaanthraquinone (dithlanon), 2-thio-1,3-dithio-[4,5-b]quinoxaline (thioquinox) methyl 1-(butylcarbamoyl)-2-benzimidazole carbamate (benomyl), 2-(4'-thiazolyl)benzimidazole (thiabendazole), 4-(2-chlorophenylhydrazono)-3-methyl-5-isoxazolone, pyridine-2-thiol-1-oxide, 8-hydroxyquinoline sulfate and metal salts thereof; 2,3-dihydro-5-carboxanilido-6-methyl-1,4-oxathin-4,4-dioxide, 2,3-dihydro-5-carboxanilido-6-methyl-1,4-oxathin, alpha-(phenyl)-alpha-(2,4-dichlorophenyl)-5-pyrimidinyl-methanol (triarimol), cis-N-[(1,1,2,2-tetrachloroethyl)thio]-4-cyclohexene-1,2-dicarboximide, 3-[2-(3,5-dimethyl-2-oxycyclohexyl)-2-hydroxy]-glutarimide (cycloheximide), dehydroacetic acid, N-(1,1,2,2-tetrachloroethylthio)-3a,4,7,7a-tetrahydrophthalimide (captfol), 5-butyl-2-ethylamino-4-hydroxy-6-methylpyrimidine (ethirimol), acetate of 4-cyclododecyl-2,6-dimethylmorpholine (dodemorph), and 6-methyl-2-oxo-1,3-dithiol[4,5-b]-quinoxaline (quinomethionate).

(d) miscellaneous halogenated fungicides such as: tetrachloro-p-benzoquinone (chloranil), 2,3-dichloro-1,4-naphthoquinone (dichlone), 1,4-dichloro-2,5-dimethoxybenzene (chloroneb), 3,5,6-trichloro-o-anisic acid (tricamba), 2,4,5,6-tetrachloroisophthalonitrile (TCPN) 2,6-dichloro-4-nitroaniline (dichloran), 2-chloro-1-nitropropane, polychloronitrobenzenes such as: pentachloronitrobenzene (PCNB) and tetrafluorodichloroacetone;

(e) fungicidal antibiotics such as:

griseofulvin, kasugamycin and streptomycin;

(f) copper-based fungicides such as:

cuprous oxide, basic cupric chloride, basic copper carbonate, copper naphthenate, and Bordeaux mixture;

(g) other rice fungicides such as:

tricyclazole, iso-prothiolane, probenazole, propiconazole, edifenphos, O-O-diisopropyl-benzyl-thiophosphate, Iprodione, procymidone, vinclozolin, benomyl, thiophanate methyl, mepronil, tencycuron and validamycin A; and

(h) miscellaneous fungicides such as:

diphenyl, dodecylguanidine acetate (dodine), phenylmercuric acetate, N-ethylmercuri-1,2,3,6-tetrahydro-3,6-endomethano-3,4,5,6,7,7-hexachlorophthalimide, phenyl mercuric monoethanol ammonium lactate, p-dimethylaminobenzenediazo sodium sulfonate, methyl isothiocyanate, 1-thiocyano-2,4-dinitrobenzene, 1-phenylthiosemicarbazide, nickel-containing compounds, calcium cyanamide, lime sulfur, sulfur, and 1,2-bis(3-methoxycarbonyl-2-thioureido)benzene (thiophanatemethyl).

The enantiomorphs, acid addition salts and metal salt complexes of this invention can be advantageously employed in various ways. Since these compounds possess broad spectrum fungicidal activity, they can be employed as fungicides in turf, fruit orchards, vegetable crops, cereal crops, golf course applications and the storage of cereal grain. Other applications of this invention will suggest themselves to those skilled in the art of agriculture and horticulture.

EXAMPLES

In summary, the substituted benzylicyanide was synthesized to the alpha-phenyl-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile in three steps. The benzylicyanide was alkylated in the first step by one of six methods. The alkylated phenylacetone nitrile was bromomethylated in the second step by one of four methods. The alkylated triazole propanenitrile was synthesized from the alkylated phenylacetone nitrile bromide by nucleophilic displacement with potassium triazole in the third step. In the third step the potassium triazole was either previously prepared and added to the bromide or equal parts of potassium hydroxide, triazole and DMSO was added to an equivalent amount of toluene, heated to 100-120°C for 2 hours to distill off the toluene and drive off the azeotropic water and the bromide added to the freshly prepared potassium triazole. In a third method the potassium triazole was prepared in situ by the reaction of potassium carbonate and triazole in a MEK or DMSO solvent.

A two step procedure can be used to prepare the alpha-phenyl-alpha-(2-phenylethyl)-1H-1,2,4-triazole-1-propanenitrile. The first step is the same as for the three step procedure. However the product of step 1 is coupled via a base with chloromethyltriazole-HCl in DMF. At least two equivalents of base is employed. The anion is generated at room temperature then the solid HCl salt is added. The second equivalent liberates the salt to the free base, chloromethyl triazole, which reacts with the anion. The base can be hydroxide, alkoxide or hydride. NaOH or KOH can be employed however NaH is preferred on an experimental scale. For less acidic intermediates where the anion is more difficult to form KH is preferred.

A similar procedure using chloromethyl triazole-HCl is to liberate the salt to the free base prior to addition. This can be done with NaOH in CH₂ Cl₂, then the solvent removed and the addition done with DMF using NaH as the base or DMSO with NaOH as the base.

The halogenated phenethyl triazoles may be prepared from the corresponding phenethyl triazoles by the reaction with n-chlorosuccinimide (NCS) or n-bromosuccinimide (NBS) in the presence of a catalyst or initiator such as benzoyl peroxide. The phenethyl triazole is dissolved in an inert solvent such as carbon tetrachloride and 1 or 2 equivalents of the halogenating agent is used.

The halogenated triazoles can be transformed into the ethenyl compound by base elimination using a metal hydroxide or alkoxide. Typical bases include sodium hydroxide, sodium methoxide and potassium tert-butoxide.

A summary of the steps and the process of preparation of the Compounds 1-96 are set forth in Tables 3 and 4.

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TABLE 3

Step 1	Step 2	Step 3
1. 50% NaOH in DMSO or DMSO/toluene (50% NaOH)	1. 50% NaOH in DMSO (50% NaOH)	1. KTriazole previously prepared (KTr)
	2. 60% NaH in DMF (NaH/DMF)	
2. 60% NaH in DMF (NaH/DMF)		2. KOHTriazole (KOHTr)
	3. 60% NaH/35% KH in DMF (NaH/KH)	
3. 60% NaH in DMF, toluene (NaH/DMFT=1:2 DMF:toluene, NaH/DMFT=2:1 DMF:toluene)	4. 35% KH in DMF (KH/DMF)	3. K_2CO_3 /Triazole (K_2CO_3 /Tr)
4. 60% NaH in DMF, benzene (NaH/DMFB)	<u>Step 2-3</u> $ClCH_2$ Triazole $(ClCH_2Tr)$	
5. 60% NaH/35% KH in DMF (NaH,KH/DMF)		
6. 60% NaH/35% KH in DMF, toluene (NaH,KH/DMFT)		

TABLE 4

<u>Compound</u>	<u>Step 1</u>	<u>Step 2</u>	<u>Step 3</u>	<u>m.p. °C</u>
1	50% NaOH	50% NaOH	KTr	113-114
2	NaH/DMF	NaH/DMF	KTr	118-119
3	NaH/DMF	50% NaOH	KTr	105-106
4	NaH/DMF	50% NaOH	KTr	100-102
5	NaH/DMF	NaH/THF	KTr	128-129
6	NaH/DMFt	50% NaOH	KOHTr	139-140
7	NaH/DMF	NaH/DMF	KTr	144-146
8	NaH/DMF	NaH/DMF	KTr	147-148
9	NaH/DMF	NaH/DMF	KTr	124-126
10	NaH/DMF	50% NaOH	KTr	122-124
11	NaH/DMF	50% NaOH	KTr	108-111
12	NaH/DMFt	50% NaOH	KTr	168-169
13	NaH/DMFt	50% NaOH	KTr	130-131
14	NaH/DMFt	50% NaOH	KOHTr	127-128
15	NaH/DMFt	50% NaOH	KOHTr	154-155
16	NaH/DMF	50% NaOH	KTr	115-117
17	NaH/DMFt	50% NaOH	KOHTr	113-114
18	NaH/DMFt	50% NaOH	KOHTr	99-102
19	NaH/DMFt	NaH/DMF	KOHTr	133-134
20	NaH/DMFB	50% NaOH	KTr	118-121
21	NaH/DMFB	50% NaOH	KTr	85-88
22	NaH/DMF	50% NaOH	KTr	142-143
23	NaH/DMFT	50% NaOH	KTr	116-119
24	NaH/DMFT	50% NaOH	KTr	119-121
25	NaH/DMF	50% NaOH	KTr	135-137
26	NaH/DMF	NaH/DMF	KTr	121-122
27	NaH/DMFT	50% NaOH	KTr	107-108
28	NaH/DMF	50% NaOH	KTr	Oil
29	NaH/DMFT	50% NaOH	KTr	Oil
30	NaH/DMF	50% NaOH	KOHTr	113-114
31	NaH/DMF	50% NaOH	KOHTr	88-90

<u>Compound</u>	<u>Step 1</u>	<u>Step 2</u>	<u>Step 3</u>	<u>m.p. °C</u>
32	NaH/DMF	NaH/DMF	KTr	119-120
33	NaH/DMF	NaH/DMF	KTr	145-146
34	NaH/DMF	NaH/DMF	KTr	95-97
35	NaH/DMF	NaH/DMF	KTr	119-120
36	NaH/DMF	NaH/DMF	KTr	119-120
37	NaH/DMF	NaH/DMF	KTr	136-137
38	NaH/DMFT	50% NaOH	K ₂ CO ₃ /Tr	110-113
39	NaH/DMFT	50% NaOH	K ₂ CO ₃ /Tr	Oil
40	NaH/DMF	ClCH ₂ Tr		102-103
41	NaH/DMF	ClCH ₂ Tr		Oil
42	NaH/DMF	ClCH ₂ Tr		122-124
43	NaH/DMF	ClCH ₂ Tr		102-103
44	NaH/DMF	ClCH ₂ Tr		110-112
45	NaH/DMF	ClCH ₂ Tr		Oil
46	NaH/DMF	ClCH ₂ Tr		79-80
47	NaH/DMF	ClCH ₂ Tr		115-117
48	NaH/DMF	ClCH ₂ Tr		121-122
49	NaH/DMF	ClCH ₂ Tr		134-135
50	NaH/DMF	ClCH ₂ Tr		Glass
51	NaH/DMF	ClCH ₂ Tr		80-81
52	NaH/DMF	ClCH ₂ Tr		112-113
53	NaH/DMF	ClCH ₂ Tr		72-73
54	NaH/DMF	ClCH ₂ Tr		118-119
55	NaH/DMF	ClCH ₂ Tr		96-97
56	NaH/DMF	ClCH ₂ Tr		109-110
57	NaH/DMF	ClCH ₂ Tr		93-94
58	NaH/DMF	ClCH ₂ Tr		Oil
59	NaH/DMFT	ClCH ₂ Tr		86-87
60	NaH/DMFT	ClCH ₂ Tr		Oil
61	NaH/DMF	NaH/DMF	KTr	111-113
62	NaH/DMF	NaH/DMF	KTr	95-97
63	NaH/DMFT	NaH-DMF	KTr	Oil
64	NaH/DMFT	NaH/DMF	KTr	98-99

<u>Compound</u>	<u>Step 1</u>	<u>Step 2</u>	<u>Step 3</u>	<u>m.p. °C</u>	
65	NaH/DMFT	NaH/DMF	KTr	108-109	
66	NaH/DMFT	NaH/DMF	KTr	108-109	
67	NaH/DMFT	NaH/DMF	KTr	112-113	5
68	NaH/DMFT	NaH/DMF	KTr	112-114	
69	NaH/DMFT	NaH/DMF	KTr	115-117	
70	NaH/DMFT	ClCH ₂ Tr		Oil	10
71	NaH/DMFT	NaH/DMF	KTr	115-116	
72	NaH/DMFT	NaH/DMF	KTr	82-83	15
73	NaH/DMFT	NaH/DMF	KTr	108-110	
74	NaH, KH/DMFT	KH/DMF	KTr	130-132	
75	NaH/DMFT	NaH/DMF	KTr	Oil	20
76	NaH/DMF	KH/DMF	KTr	96-98	
77	NaH/DMF	NaH/KH	KTr	80	
78	NaH/DMF	NaH/KH	KTr	126-128	25
79	NaH/DMF	NaH/KH	KTr	97-98	
80	NaH/DMFT	NaH/DMF	KTr	144-146	
81	NaH/DMFT	NaH/DMF	KTr	104-106	30
82	NaH/DMFT	NaH/DMF	KTr	98-100	
83	NaH/DMFT	ClCH ₂ Tr		100-101	
84	NaH/DMFT	ClCH ₂ Tr		103-104	35
85	NaH/DMFT	ClCH ₂ Tr		96-98	
86	NaH/DMFT	ClCH ₂ Tr		126-128	
87	NaH/DMFT	ClCH ₂ Tr		Oil	40
88	NaH, KH/DMF	ClCH ₂ Tr		Oil	
89	NaH/DMFT	ClCH ₂ Tr		115-117	
90	NaH/DMFT	ClCH ₂ Tr		Oil	45
91	NaH/DMFT	ClCH ₂ Tr		108-110	
92	NaH/DMFT	ClCH ₂ Tr		130-131	
93	50% NaOH	50% NaOH	K ₂ CO ₃ /Tr	91-94	50
94	50% NaOH	NaH/DMF	KTr	244-248	
95	NaH/DMFT	NaH/DMF	KTr	Oil	
96	NaH/DMF	ClCH ₂ Tr		Oil	55

60

The melting points and elemental analysis for Compounds 1-96 are set forth in Tables 4 and 5. The amount of chlorine, fluorine and oxygen were not measured in all of the examples. The NMR for Compounds 70, 95 and 96

65

follow Table 5.

TABLE 5
Elemental Analysis

Com- pound	Carbon		Hydrogen		Nitrogen		Oxygen		Chlorine		Bromine		Fluorine	
	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found
1	67.75	67.73	5.09	5.06	16.64	16.49			10.52	10.47				
2	75.47	75.97	6.00	6.05	18.53	18.34								
3	72.27	72.54	6.06	6.24	16.86	16.25	4.81	5.66						
4	71.23	71.51	5.35	5.35	17.49	16.83							5.93	5.87
5	61.44	62.04	4.34	4.49	15.07	14.98								
6	61.45	62.06	4.34	4.32	15.89	13.52			19.10	18.91				
7	64.29	63.94	4.54	4.77	15.58	15.79			19.11	19.79				
8	68.47	68.29	5.46	5.58	15.97	16.34			9.98	9.86			5.31	5.52
9	65.49	65.49	5.23	5.42	15.30	15.29	4.40	4.49	10.10	10.04				
10	67.73	68.37	5.09	5.17	16.64	16.29			9.70	9.98			5.93	5.54
11	71.21	72.00	5.35	5.36	17.50	17.14			10.53	10.15				
12	79.32	78.15	5.86	5.69	14.81	14.09								
13	64.83	64.35	4.63	4.62	15.14	14.86							15.39	15.58
14	64.83	65.13	4.63	4.53	15.14	15.09							15.39	15.22
15	61.44	61.30	4.34	4.38	15.10	14.86			19.11	19.00				
16	59.83	61.68	4.49	4.67	14.70	14.28					20.96	19.10		

Com- pound	Carbon		Hydrogen		Nitrogen		Oxygen		Chlorine		Bromine		Fluorine	
	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found
17	67.73	67.71	5.09	5.00	16.64	16.41			10.53	10.98				
18	67.73	67.69	5.09	5.21	16.64	16.41			10.53	10.79				
19	64.63	64.64	4.63	4.84	15.74	14.89							15.39	15.17
20	59.32	59.46	3.99	4.20	13.86	13.25			8.76	9.48			14.09	13.36
21	61.83	62.65	4.16	4.24	14.41	13.71							19.57	17.89
22	56.87	57.98	3.88	3.88	13.48	13.06			8.53	9.73	19.23	18.87		
23	59.32	59.11	3.99	4.00	13.85	13.20			8.76	9.16			14.09	13.65
24	61.83	61.85	4.16	4.31	14.44	13.61							19.67	19.36
25	57.13	57.80	4.04	4.05	14.04	13.69					20.02	19.57	4.76	4.78
26	64.29	64.23	4.55	4.65	15.80	15.45			9.99	9.54			5.36	5.30
27	59.83	60.85	4.49	4.27	14.70	14.28					20.96	19.47		
28	67.42	68.26	4.76	4.79	16.56	16.33							11.23	10.78
29	62.99	63.16	4.78	5.07	13.99	13.62	4.01						14.23	14.37
30	64.29	64.35	4.55	4.54	15.80	15.74			9.99	10.09			5.86	
31	61.94	61.83	4.35	4.53	25.20	24.93			19.10	19.28				
32	59.83	59.92	4.49	4.64	14.70	14.16							20.96	21.43
33	57.13	57.32	4.04	4.34	14.04	13.66					20.02	20.16	4.78	4.60
34	67.13	66.66	5.09	5.07	16.64	16.02			10.53	10.56				
35	61.44	61.42	4.34	4.47	15.10	14.75			19.11	19.05				
36	63.75	63.85	5.35	4.73	15.66	15.32			9.91	9.90			5.31	4.83

Com- pound	Carbon		Hydrogen		Nitrogen		Oxygen		Chlorine		Bromine		Fluorine	
	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found
37	54.87	55.20	3.88	4.08	13.48	12.44			8.53	7.70	19.23	20.31		
38	57.13	57.37	4.04	4.25	14.04	14.25					20.02	20.05	4.76	
39	54.87	54.40	3.88	4.48	13.48	11.88			8.53	8.60	19.23	17.77		
40	61.44	61.19	4.35	4.32	15.10	14.95			19.10	18.88				
41	64.29	61.57	4.55	4.58	15.80	14.88			10.00	10.44			5.36	4.86
42	64.29	63.73	4.55	4.53	15.79	15.46			10.00	9.89			5.35	5.37
43	67.42	67.14	4.77	4.82	16.56	16.46							11.24	11.16
44	64.30	63.82	4.55	4.56	15.80	15.84			10.00	9.87			5.36	5.30
45	66.52	67.42	4.77	5.00	16.57	16.56							11.24	11.39
46	59.31	59.58	3.99	4.08	13.84	13.65			8.76	8.58			14.09	14.02
47	64.29	64.12	4.55	4.52	15.80	15.27			9.99	10.19			5.36	4.97
48	54.87	54.80	3.88	4.24	13.48	13.30			8.53	8.50	19.23	19.17		
49	65.46	65.54	5.22	5.23	15.28	14.61	4.36	5.14	9.67	9.44				
50	64.83	65.02	4.63	4.62	15.14	15.13							15.40	15.58
51	57.52	57.53	3.68	3.74	12.79	12.77							26.02	25.85
52	71.21	70.96	5.35	5.47	17.50	17.51							5.93	5.82
53	59.31	59.65	4.15	4.26	14.43	13.94			8.76				14.09	13.87
54	59.83	59.92	4.49	4.61	14.70	14.68					20.97	20.77		
55	53.45	53.05	3.59	3.54	12.48	12.29					17.79	17.68	12.70	12.59
56	61.83	61.92	4.15	4.26	14.43	14.48							19.58	19.47

Com- pound	Carbon		Hydrogen		Nitrogen		Oxygen		Chlorine		Bromine		Fluorine	
	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found
57	59.31	59.68	3.99	4.17	13.84	13.76			8.76	8.80			14.08	13.86
58	72.25	72.21	6.07	6.61	16.87	15.66	4.82	6.20						
59	72.25	72.53	6.07	5.89	16.87	15.79	4.82	5.43						
60	69.57	68.93	6.12	6.43	15.47	15.03	8.83	9.85						
61	65.46	65.87	5.22	5.34	15.28	14.82	4.36	4.68	9.67	9.59				
62	72.25	72.25	6.07	6.21	16.96	16.62	4.81	5.09						
63	64.30	64.73	4.55	4.86	15.80	14.66			9.99	9.93			5.36	4.80
64	58.60	58.43	3.89	3.91	14.40	14.43			18.22	17.89			4.88	4.78
65	56.22	56.49	3.73	3.79	13.82	13.79			26.23	26.17				
66	54.87	54.87	3.88	3.90	13.48	13.56			8.53	8.52	19.23	18.77		
67	54.87	54.79	3.88	3.90	13.48	13.47			8.53	8.85	19.23	18.83		
68	57.13	57.03	4.04	4.09	14.04	14.14					20.02	19.46	4.76	4.94
69	57.13	57.04	4.04	4.08	14.04	14.04					20.02	19.48	4.76	4.81
70	See NMR													
71	67.42	65.20	4.77	5.01	16.59	14.56							11.24	11.09
72	61.83	61.92	4.16	4.15	14.43	14.36							19.58	19.81
73	78.36	78.49	5.72	5.74	15.91	15.81								
74	59.28	59.42	4.98	5.06	13.18	13.12	3.76	3.99			18.80	18.60		
75	53.45	53.21	3.59	3.78	12.48	12.46					17.79	17.14	12.69	12.03
76	63.74	62.62	5.11	5.28	13.76	13.41	3.86						13.76	13.68

Compound	Carbon		Hydrogen		Nitrogen		Oxygen		Chlorine		Bromine		Fluorine	
	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found	Calc.	Found
77	72.79	73.19	6.41	6.40	16.18	16.12	4.62	4.80						
78	66.09	66.31	5.26	5.37	13.76	13.73	4.64	4.21	10.00	9.92				
79	68.79	68.89	5.77	5.67	15.49	15.31	4.80	4.73					5.36	5.40
80	61.44	60.53	4.34	4.07	15.10	14.60			19.11	18.77				
81	61.44	61.16	4.34	4.17	15.10	15.18			19.11	19.36				
82	64.30	64.32	4.55	4.15	15.80	16.02			10.00	10.23			5.36	5.16
83	71.21	71.20	5.35	5.36	17.50	17.44							5.93	5.93
84	65.46	65.53	5.22	5.25	15.28	15.23	4.36	4.34	9.67	9.22				
85	54.87	54.81	3.88	3.78	13.48	13.34			8.53	8.00	19.23	18.90		
86	64.30	64.35	4.55	4.85	15.80	15.64			10.00	9.99			5.36	5.15
87	59.83	58.56	4.50	4.51	14.70	13.56					20.96	20.91		
88	59.28	58.40	4.98	5.43	13.18	11.56	3.76				18.80	17.69		
89	67.43	67.87	4.77	4.89	16.59	16.46							11.24	11.54
90	49.57	49.69	3.51	3.31	12.18	11.97					34.74	34.61		
91	64.31	64.10	4.51	4.62	15.80	15.52			10.01	9.96			5.36	5.31
92	61.45	61.54	4.31	4.52	15.09	15.06			19.14	19.08				
93	72.24	72.55	6.06	6.39	16.87	16.09	4.82	5.21						
94	72.73	72.03	5.09	5.12	13.58	12.31			8.61	8.72				
95	See NMR													
96	See NMR													

NMR was measured for Compounds 70, 95 and 96. The results are as follows:

Compound 70: NMR(90 MHz): (Mixture of diastereoisomers): 1.2-1.6(two doublets, 3H), 2.4-2.8(m, 3H),

4.6-4.9(two overlapping ABq, 2H), 7.0-7.8(m, 10H), 7.90(s, 1H) and 8.0(s, 1H).

Compound 95: NMR(90 MHz): 2.0-2.4(m, 4H), 4.8(ABq, 2H), 6.9-7.4(m, 7H), 7.8(s, 1H) and 7.9(s, 1H).

Compound 96: NMR(90 MHz): 2.2-2.9(m, 4H), 4.7-5.0(ABq, 2H), 7.0-7.4(ABq, 4H), 7.4-7.8(m, 3H) and 8.6-8.7(br d, 1H).

The following are examples of the preparation of typical compounds of the present invention.

PREPARATION OF COMPOUND

14 - Alpha-phenyl-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of alpha-[2-(3-trifluoromethylphenyl)ethyl]phenylacetonitrile

A 500 ml 4 neck round bottom flask was charged with 11.0 gms of 60% NaH (0.275 mole, 1.1 eq.), washed twice with hexane, in 50 ml of dry DMF. The reaction was cooled to 0°C and 29.4 gms of benzyl cyanide (0.25 mole, 1.0 eq.) in 50 ml of toluene was added dropwise. The reaction was warmed to room temperature and then cooled to -20°C and 52 gms (0.25 mole, 1.0 eq.) of 3-trifluoromethylphenethyl chloride in 50 ml of DMF was added dropwise. The reaction was stirred at -20°C for 3 hours after which gas liquid chromatography showed 60% monoalkylation and 40% dialkylation. The reaction was quenched with water and extracted with ether. After drying and concentrating, the crude product was chromatographed by high pressure liquid chromatography (95:5) hexane:ethyl acetate and gave 19.2 gms of 97% pure product (26.3% yield). The mixture was added directly in the next step.

NMR (60 MHz): 2.0-2.4(m, 2H), 2.7-3.0(m, 2H), 3.6-3.9(t, 1H) and 7.2-7.4(d, 9H).

Step 2 - Preparation of 1-bromo-2-cyano-2-phenyl-4-(3-trifluoromethylphenyl)butane

A 500 ml 4 neck flask was charged with 19.0 gms of alpha-[2-(3-trifluoromethylphenyl)ethyl]phenyl acetonitrile (0.064 mole, 1.0 eq.) and 17 gms of CH₂Br₂ (0.097 mole, 1.5 eq.) in 50 ml of DMSO. The reaction was stirred at room temperature and 10.3 gms of 50% NaOH (0.128 mole, 2.0 eq.) was added dropwise. The reaction was warmed to 50°C and stirred for 2 hours. The reaction was worked up with water and ether and gave after drying and concentrating 25 gms of product which was used directly in the triazole coupling (96.9% yield).

NMR (60 MHz): 2.2-2.8(m, 4H), 3.7(s, 2H) and 7.2-7.6(m, 9H).

Step 3 - Preparation of alpha-phenyl-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1,2,4-triazole-1-propanenitrile

A 250 ml 4 neck round bottom flask was charged with 4.05 gms of 87% KOH (0.062 mole, 1.25 eq.) and 4.8 gms (0.068 mole, 2.2 eq.) of triazole in 25 ml DMSO. The reaction was warmed to 90°C until homogeneous and 25 ml of toluene was added and azeotroped for 4 hours. The toluene was distilled off at 165°C and the reaction was cooled to 100°C and 12.5 g of 1-bromo-2-cyano-2-phenyl-4-(3-trifluoromethylphenyl)butane (0.031 mole, 1.0 eq.) was added. The reaction was heated at 125°C for 1 hour and then worked up with water and ethyl acetate, dried and concentrated. Purification by high pressure liquid chromatography (1:1) hexane:ethyl acetate gave 8.8 gms of a white solid having a melting point of 127-128°C (76.7% yield).

IR (nujol, cm⁻¹): 2980(s), 2240(w), 1430(s), 1380(s), 1330(s), 1280(m), 1200(m), 1160(s), 1140(s), 1130(s), 1120(s), 1075(m), 1025(w), 800(m), 710(s) and 670(s).

NMR (60 MHz): 2.4-2.9(m, 4H), 4.6(s, 2H), 7.3-7.5(s, 4H), 7.7(s, 1H) and 7.9(s, 1H).

Elemental Analysis: C₂₀ H₁₇ N₄ F₃

Theor: C: 64.83, H: 4.83, N: 15.14, F: 15.39

Found: C: 65.13, H: 4.52, N: 15.09, F: 15.22

PREPARATION OF COMPOUND

20 - Alpha-(4-chlorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of alpha-[2-(3-trifluoromethylphenyl)ethyl]-4-chlorophenylacetonitrile

A 1 neck 500 ml round bottom flask was charged with 4.4 gms of 60% NaH (0.11 mole, 1.0 eq.), washed three times with 25 ml hexanes, in 100 ml of 2:1 benzene/DMF. Then 15.1 gms (0.10 mole, 1.0 eq.) of 4-chlorobenzyl cyanide was added and stirred for 2 hours at room temperature. While stirring at room temperature 20.8 gms (0.1 mole, 1.0 eq.) of 3-(trifluoromethyl)phenethyl chloride was added dropwise over several hours and then stirred at room temperature overnight. The product was worked up with water and ether and distilled after concentrating to give 16.2 gms of product (50.1% yield).

NMR (60 MHz): 2.0-3.0(m, 4H), 3.5-3.8(t, 1H), 7.3(s, 4H) and 7.5(s, 4H).

Step 2 - Preparation of 1-bromo-2-cyano-2-(4-chlorophenyl)-4-(3-trifluoromethylphenyl)butane

A 1 neck 300 ml round bottom flask was charged with 16.2 gms (0.05 mole, 1.0 eq.) of alpha-[2-(3-trifluoromethylphenyl)ethyl]-4-chlorophenylacetonitrile and 17.4 gms of CH₂Br₂ (0.10 mole, 2.0 eq.) in 50 ml of DMSO. The reaction was stirred at room temperature and 10 gms of 50% NaOH was added dropwise and an exotherm was observed. The reaction was stirred for 45 minutes and gas liquid chromatography showed the reaction was complete. The product was worked up with ether and water. Drying and removal of the solvent gave 20.8 gms of a yellow oil (100% yield).

NMR (60 MHz): 2.0-3.2(m, 4H), 3.8(s, 2H) and 7.5-7.6(d, 10H).

Step 3 - Preparation of

alpha-(4-chlorophenyl)-alpha-[2-(3-trifluoromethylphenyl)ethyl]-1,2,4-triazole-1-propanenitrile

A 500 ml 1 neck round bottom flask stirred under nitrogen equipped with condenser and addition funnel was charged with 10.7 gms of potassium triazole (0.10 mole, 4.0 eq.), 75 ml of DMSO and 1-bromo-2-cyano-2-(4-chlorophenyl)-4-(3-trifluoromethylphenyl)butane (10.4 gms, 0.025 mole, 1.0 eq.). The reaction was heated at 80°C overnight and then quenched by adding 1 liter of H₂O and extracted three times with 200 ml of ether. The combined ether extracts were washed with water and brine, then dried and rotovaped to give a crude product which was slurried in ether:hexane (1:1). A solid formed which was washed with hexane and gave 5.0 gms of a light yellow solid having a melting point of 118-121°C (47% yield).

NMR (60 MHz): 2.4-3.2(m, 4H), 5.0-5.2(br s, 2H), 7.6-7.9(br s, 8H), 8.1(s, 1H) and 8.4 (s, 1H).

Elemental Analysis: C₂₀ H₁₆ N₄ F₃ Cl

Theor: C: 59.32, H: 3.99, N: 13.85, Cl: 8.76, F: 14.09

Found: C: 59.46, H: 4.20, N: 13.25, Cl: 9.48, F: 13.36

PREPARATION OF COMPOUND

44 - Alpha-(2-chlorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of 2-(4-fluorophenyl)ethyl methanesulfonate. Mesylation of 2-(4-fluorophenyl)ethanol.

A 500 ml 3 neck round bottom flask stirred under nitrogen equipped with condenser and addition funnel was charged with 42.04 gms of 2-(4-fluorophenyl)ethanol (0.3 mole, 1.0 eq.) in 100 ml of tetrahydrofuran (THF). The reaction was cooled to 10°C and 60.7 gms (0.60 mole, 2.0 eq.) of triethylamine was added directly. This was followed by dropwise addition of 68.73 grams (0.6 mole, 2.0 eq.) of methanesulfonyl chloride in 30 ml of THF maintaining the temperature below 30°C. An additional 150 ml of THF was added and the reaction stirred for 6 hours. The reaction was quenched with 200 ml of water and 300 ml of ether was added. The ether was washed with 75 ml of 10% HCl, twice with 50 ml of saturated NaHCO₃, twice with 50 ml of water, dried and concentrated to give 58.0 gms of a brown liquid (88.7% yield).

NMR (90 MHz): 2.9(s, 3H), 2.9-3.2(m, 2H), 4.2-4.4(t, 2H) and 6.9-7.4(m, 4H).

Step 2 - Preparation of alpha-[2-(4-fluorophenyl)ethyl]-2-chlorophenylacetonitrile

A 4 neck 300 ml round bottom flask was charged with 4.26 gms of 60% NaH (0.105 mole, 1.05 eq.), washed three times with 25 ml of hexanes, in 60 ml of DMF. Then 15.16 gms (0.10 mole, 1.0 eq.) of 2-chlorophenyl acetonitrile in 40 ml of DMF was added and stirred for 1 hour at 10°C. Then 2-(4-fluorophenyl)ethyl methanesulfonate (22.2 gms, 0.102 mole) in 50 ml of DMF was added dropwise. The reaction was complete after 2 hours, then quenched with 10 ml of 10% HCl. Then 60 ml of water was added and extracted with 200 ml of ether which was then washed twice with 50 ml of 10% HCl, then dried and concentrated. 27.15 gms of crude product resulted which was distilled under reduced pressure and gave 17.46 gms of an oil having a boiling point of 180-188°C at 1 mm of Hg (63.5% yield).

NMR (90 MHz): 2.2-2.4(t, 2H), 2.8-3.0(m, 2H), 3.8-3.9(t, 2H) and 6.9-7.5(m, 8H).

Step 3 - Preparation of alpha-(2-chlorophenyl)-alpha-[2-(4-fluorophenyl)ethyl]-1,2,4-triazole-1-propanenitrile

A 200 ml 3 neck round bottom flask was charged with 2.3 gms of 60% NaH (0.055 mole, 2.75 eq.), washed twice with 25 ml hexanes, in 40 ml of DMF. The reaction was cooled to 10°C and 5.46 gms (0.02 mole, 1.0 eq.) of alpha-[2-(4-fluorophenyl)ethyl]-2-chlorophenylacetonitrile in 40 ml of DMF was added dropwise over ten minutes. After 20 minutes, 3.12 gms (0.0204 mole, 1.02 eq.) of chloromethyltriazole-HCl was added directly in two portions. After 1 hour, gas liquid chromatography indicated the reaction was complete and the reaction was quenched by slowly adding 20 ml of water. The product was extracted with 200 ml of CH₂Cl₂ and washed twice with 50 ml of water. After drying and concentrating, an orange oil was obtained which crystallized from ethyl ether. The product was filtered and gave 2.5 gms of a light tan solid having a melting point of 110-112°C (35% yield).

IR (nujol, cm⁻¹): 1505(m), 1440(s), 1370(m), 1270(m), 1220(m), 1130(m) and 750(m).

NMR (90 MHz): 2.2-3.0(m, 4H), 3.8-4.2(ABq, 2H), 6.9-7.4(m, 7H), 7.8(s, 1H) and 7.9(s, 1H).

Elemental Analysis: C₁₈ H₁₆ N₄ FCl

Theor: C: 64.30, H: 4.55, N: 15.80, F: 5.36, Cl: 10.00

Found: C: 63.82, H: 4.56, N: 15.84, F: 5.30, Cl: 9.87

PREPARATION OF COMPOUND

64 - Alpha-(2-chloro-6-fluorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of alpha-[2-(4-chlorophenyl)ethyl]-2-chloro-6-fluorophenylacetonitrile

A 3 neck 200 ml round bottom flask was charged with 7.5 gms of 60% NaH (0.187 mole, 1.5 eq.), washed three times with 25 ml hexanes, in 60 ml of 2:1 toluene:DMF. To this was added 21.2 gms (0.125 mole, 1.0 eq.) of 2-chloro-6-fluorophenylacetonitrile dropwise over 0.5 hour in 40 ml of 2:1 toluene:DMF. The reaction was stirred for 20 minutes at 10°C then at room temperature for 1 hour after which 32.1 gms of 2-(4-chlorophenyl)ethyl methanesulfonate (0.137 mole, 1.1 eq.) in 60 ml of 2:1 toluene:DMF was added

dropwise over 1 hour. Approximately 70 ml of 2:1 toluene:DMF was added to permit constant stirring and the reaction was stirred for an additional 3.5 hours after which gas liquid chromatography indicated the reaction was complete. Then 50 ml of water was added, followed by 10 ml of 10% HCl and 300 ml of ether. The ether was washed with 100 ml of water which was extracted twice with 50 ml of ether then washed with water. The combined ethers were dried and concentrated to give 40.0 gms of crude product which was distilled under reduced pressure. 26.6 gms (69.8%) of product resulted having a boiling point of 175-185°C at 1 mm Hg. NMR (90 MHz): 2.2-2.9(m, 4H), 4.2-4.4(t, 1H) 7.0-7.4(m, 7H).

Step 2 - Preparation of 1-bromo-2-cyano-2-(2-chloro-6-fluorophenyl)-4-(4-chlorophenyl)butane

A 3 neck 200 ml round bottom flask was charged with 2.4 gms of 60% NaH (0.048 mole, 1.2 eq.), washed twice with 25 ml of hexanes, in 40 ml of DMF. At room temperature 12.28 gms (0.04 mole, 1.0 eq.) of alpha-[2-(4-chlorophenyl)ethyl]-2-chloro-6-fluorophenyl-acetonitrile was added dropwise over 0.5 hour in 30 ml of DMF. Then 10.43 gms of CH₂ Br₂ (0.60 mole, 1.5 eq.) in 20 ml of DMF was added dropwise and the reaction was stirred at room temperature for 0.5 hour after which gas liquid chromatography indicated that the reaction was complete. The reaction was quenched after 1 hour by adding 20 ml of water and 200 ml of ether which was separated. After washing with water, drying and concentrating 14.84 gms (92.9% yield) resulted which was used directly in the triazole coupling.

NMR (90 MHz): 2.4-2.8(m, 4H), 3.8-4.2(ABq, 2H) and 6.9-7.3(m, 7H).

Step 3 - Preparation of

alpha-[2-(2-chloro-6-fluorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1,2,4-triazole-1-propanenitrile

A 200 ml 1 neck round bottom flask was charged with 14.84 gms (0.038 mole, 1.0 eq.) of 1-bromo-2-cyano-2-(2-chloro-6-fluorophenyl)-4-(4-chlorophenyl)butane in 50 ml of DMSO. To the reaction was added 4.89 gms (0.0457 mole) of Ktriazole in 30 ml of DMSO and the flask was heated to 120°C. After 1.5 hours at 120°C, gas liquid chromatography indicated that the reaction was complete and was quenched by cooling to room temperature and adding 50 ml of water and 250 ml of ethyl acetate. After separating, the organic phase was washed twice with 100 ml of water, concentrated and triturated with ether. The solid was filtered and the filtrate concentrated and triturated with ether. The additional solid was filtered and a total of 8.06 gms was obtained having a melting point of 98-99°C (54.6% yield).

IR (nujol, cm⁻¹): 3020(m), 1600(s), 1570(m), 1500(s), 1450(s), 1275(s), 1240(m), 1205(m), 1135(m), 910(m), 890(s) and 790(s).

NMR (90 MHz): 2.4-3.2(m, 4H), 4.8-5.2(ABq, 2H), 6.9-7.4(m, 7H), 7.9(s, 1H) and 8.2(s, 1H).

Elemental Analysis: C₁₉ H₁₅ N₄ FCl₂

Theor: C: 58.60, H: 3.89, N: 14.40, F: 4.88, Cl: 18.22

Found: C: 58.43, H: 3.91, N: 14.43, F: 4.78, Cl: 17.89

PREPARATION OF COMPOUND

74 - Alpha-[2-(4-bromophenyl)ethyl]-alpha-[2-(ethoxyphenyl)-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of alpha-[2-(4-bromophenyl)ethyl]-2-ethoxyphenylacetonitrile

A 4 neck 1 liter round bottom flask was charged with 5.8 gms of 60% NaH (0.14 mole, 1.4 eq.), washed three times with 25 ml hexanes, in 100 ml of 2:1 toluene:DMF. Then 16.1 gms (0.10 mole, 1.0 eq.) of 2-ethoxybenzylcyanide in 200 ml of 2:1 toluene:DMF was added and stirred at room temperature for 2 hours. After this time, 5.7 gms of KH (0.05 mole, 0.5 eq.) washed with 25 ml of hexanes, in 50 ml of 2:1 toluene:DMF was added. After an additional 1 hour, 31 gms (0.11 mole, 1.1 eq.) of 4-bromophenethyl methane sulfonate in 100 ml of 2:1 toluene:DMF was added dropwise over 10 minutes. The reaction was stirred overnight at room temperature. The reaction was 81% monoalkylated product as indicated by gas liquid chromatography. The reaction was quenched with 10% HCl and 400 ml of ether was added and washed four times with 150 ml of water. The solvent was dried over magnesium sulfate then concentrated and gave 37 gms of a crude orange oil from which a solid crystallized out and was filtered and washed with cold toluene. 14.8 gms of a white solid resulted (43.7% yield).

NMR (90 MHz): 1.3-1.5(t, 3H), 2.0-2.3(m, 2H), 2.2-2.9(m, 2H), 3.9-4.2(m, 3H) and 6.8-7.5(m, 8H).

Step 2 - Preparation of 1-bromo-2-cyano-2-(2-ethoxyphenyl)-4-(4-bromophenyl)butane

A 3 neck 500 ml round bottom flask was charged with 3.5 gms of 100% KH (0.087 mole, 2.0 eq.) washed twice with 25 ml of hexanes, in 25 ml of DMF. At room temperature 14.8 gms (0.044 mole, 1.0 eq.) of alpha-[2-(4-bromophenyl)ethyl]-2-ethoxyphenylacetonitrile in 60 ml of DMF was added dropwise. After 1 hour, CH₂ Br₂ (0.066 mole, 1.5 eq.) in 40 ml of DMF was added dropwise. The reaction exothermed to 45°C and was stirred at room temperature for 3 hours after which gas liquid chromatography indicated the reaction was 85% complete. The reaction was stirred overnight at room temperature and the 0.8 gms of 100% KH (0.022 mole, 0.5 eq.) in 20 ml of DMF was added followed by 1.2 gms of CH₂ Br₂ in 5 ml of DMF. After 1 hour, the reaction was complete. The reaction was quenched by adding 75 ml of 10% HCl followed by addition of 300 ml of ether. After washing four times with 100 ml water, the product was dried and the solvent removed at the rotovap. 18.3 gms of an orange yellow oil resulted (96.8% yield).

NMR (60 MHz): 1.3-1.6(t, 3H), 2.2-2.8(m, 4H), 3.7-4.2(m, 4H) and 6.8-7.6(m, 8H).

Step 3 - Preparation of alpha-[2-(4-bromophenyl)ethyl]-alpha-(2-ethoxyphenyl)-1,2,4-triazole-1-propanenitrile

A 300 ml 3 neck round bottom flask was charged with 18.3 gms of 1-bromo-2-cyano-2-(2-ethoxyphenyl)-4-(4-bromophenyl)butane (0.43 mole, 1.0 eq.) in 100 ml of DMSO. To the reaction was added at room temperature 5.4 gms of KTriazole (0.050 mole) and the reaction was stirred at 100°C for 20 hours. Gas liquid chromatography indicated the reaction was complete and the reaction was quenched by adding 10% HCl and ether. The product crystallized out during the extraction and the solvent was reduced in volume and the product was filtered. 11.3 gms of a light tan solid having a melting point of 130-132°C was isolated (63% yield).

IR (nujol, cm^{-1}): 1590(w), 1265(m), 1245(m), 1135(m), 1030(m) and 750(m).

NMR (60 MHz): 1.4-1.6(t, 3H), 1.9-2.9(m, 4H), 4.0-4.4(q, 2H), 4.9(s, 2H), 6.8-7.4(m, 8H) and 7.8(s, 2H).

Elemental Analysis: $\text{C}_{21} \text{H}_{21} \text{N}_4 \text{BrO}$

Theor: C: 59.28, H: 4.98, N: 13.18, O: 3.76, Br: 18.80

Found: C: 59.42, H: 5.06, N: 13.12, O: 3.99, Br: 18.60

PREPARATION OF COMPOUND83 - Alpha-[2-(3-fluorophenyl)ethyl]-alpha-phenyl-1,2,4-triazole-1-propanenitrileStep 1 - Preparation of 1-(hydroxymethyl)-1,2,4-triazole

A 3 neck 500 ml round bottom flask equipped with condenser and mechanical stirrer was charged with 69.1 gms of triazole (1 mole), 30.1 gms of paraformaldehyde and 1 ml of triethylamine in 250 ml of THF. The reaction was stirred under nitrogen at reflux for 18 hours after which the mixture was concentrated on the rotovap. A white solid resulted which was filtered and washed with ether and gave 96.8 gms of product (97.6% yield) having a melting point of 67-70°C. The product can be further purified by dissolving in hot acetone, cooling to room temperature, filtering the solid and washing with ether.

Step 2 - Preparation of 1-(chloromethyl)-1,2,4-triazole hydrochloride

A 4 neck 1 liter flask equipped with condenser, addition funnel and mechanical stirrer was charged 45 gms of 1-(hydroxymethyl)-1,2,4-triazole (0.464 mole) in 500 ml of THF was heating to 40°C with vigorous stirring. Then SOCl_2 (61 ml, 0.84 mole) was added dropwise maintaining the temperature at 45°C. During the addition, a precipitate formed and the mixture was stirred for an additional 2 hours. The product was filtered, washed three times with ethyl acetate and vacuum dried at room temperature. 67.3 gms of product having a melting point of 118-130°C resulted (94.2% yield).

Step 3 - Preparation of 2-(3-fluorophenyl)ethanol. Reduction of 3-fluorophenylacetic acid via diborane reduction.

A 2 liter 4 neck flask stirred under nitrogen, equipped with condenser and addition funnel, was charged with 75 gms of 3-fluorophenylacetic acid (0.48 mole) in 100 ml of THF. Then 500 ml of 1M diborane THF complex (0.50 mole) was added dropwise. Upon addition, gas evolved and the reaction was cooled to maintain the temperature below 10°C. After the addition was complete, the reaction was stirred at room temperature until, by thin layer chromatography, the reaction was complete. The reaction was quenched by adding ice water and the product was extracted into ether then washed with 5% NaOH, 5% HCl, water and dried over magnesium sulfate. After concentration, 77.3 gms of a brown oil resulted.

NMR (90 MHz): 2.6(2, 1H), 2.7-2.9(t, 2H), 3.7-3.9(t, 2H) and 6.9-7.4(m, 4H).

Step 4 - Preparation of 2-(3-fluorophenyl)ethyl chloride. Chlorination of 2-(3-fluorophenyl)ethanol.

A 500 ml 4 neck round bottom flask stirred under nitrogen, equipped with condenser and addition funnel, was charged with 14.0 gms of 2-(3-fluorophenyl)ethanol (0.1 mole, 1.0 eq.) in 60 ml of toluene. Then 22 ml of SOCl_2 (35.9 gms, 0.30 mole, 3.0 eq.) was added dropwise while maintaining the temperature below 15°C with external cooling. Pyridine (8.7 gms, 1.1 eq.) was added dropwise in 10 ml of toluene. The reaction was stirred overnight at room temperature after which thin layer chromatography indicated the reaction was complete. The mixture was concentrated and the product isolated after adding water, extracting with ether and washing with water. After drying and concentrating, 10.2 gms of an orange oil resulted (64.5% yield).

NMR (60 MHz): 2.9-3.2(m, 2H), 3.5-3.8(m, 2H) and 6.8-7.4(m, 4H).

Step 5 - Preparation of alpha-[2-(3-fluorophenyl)ethyl]-phenylacetoneitrile

A 3 neck 1 liter round bottom flask was charged with 5.8 gms of 60% NaH (0.140 mole, 1.4 eq.), washed three times with 25 ml hexanes, in 100 ml of 2:1 toluene:DMF. Then 11.7 gms (0.10 mole, 1.0 eq.) of benzyl cyanide in 150 ml of 2:1 toluene:DMF was added and stirred for 2 hours. Then 2-(3-fluorophenyl)ethyl chloride (20 gms, 0.126 mole) in 100 ml of 2:1 toluene:DMF was added. The reaction was complete after 3 hours, then quenched with 10 ml of 10% HCl. Water was added and extracted with 300 ml of ether which was washed four times with 100 ml of water, then dried and concentrated. 22.5 gms of crude product resulted which was distilled under reduced pressure. 11.2 gms of product having a boiling point of 160-163°C at 1 mm Hg resulted (46.9% yield).

NMR (90 MHz): 2.0-2.3(m, 2H), 2.6-2.8(m, 2H), 3.6-3.8(t, 1H) and 6.8-7.5(m, 9H).

Step 6 - Preparation of alpha-[2-(3-fluorophenyl)ethyl]-alpha-phenyl-1,2,4-triazole-1-propanenitrile

A 500 ml 4 neck round bottom flask was charged with 0.3 gms of 60% NaH (0.075 mole, 3.0 eq.), washed twice with 25 ml of hexanes, in 50 ml of DMF. To the reaction was added dropwise 6.0 gms of alpha-[2-(3-fluorophenyl)ethyl]-phenylacetonitrile (0.025 mole, 1.0 eq.) in 200 ml of DMF. After 1 hour, 6.0 gms of chloromethyltriazole-HCl (0.038 mole, 1.5 eq.) was added directly in two portions. After 1 hour, gas liquid chromatography indicated the reaction was incomplete and an additional 1.5 gms (1.5 eq.) of 60% NaH was added in 25 ml of DMF after washing with 25 ml of hexanes. The reaction was complete in 1 hour and then quenched by adding ether containing methanol. Then 25 ml of 10% HCl was added, followed by 300 ml of ether. Washing four times with water (150 ml) was followed by drying and concentrating to give 5 gms of a yellow oil. The product crystallized out of ether and gave 2.3 gms of product having a melting point of 100-101°C (29% yield).

IR (nujol, cm^{-1}): 1580(w), 1265(m) and 1140(m).

NMR (90 MHz): 2.2-2.8 (m, 4H), 4.6(d, 2H), 6.8-7.6(m, 9H) and 7.8-7.9(d, 2H).

Elemental Analysis: $\text{C}_{19} \text{H}_{17} \text{N}_4 \text{F}$

Theor: C: 71.21, H: 5.35, N: 5.94, F: 17.50

Found: C: 71.20, H: 5.36, N: 5.93, F: 17.44

PREPARATION OF COMPOUND 88 - Alpha-[2-(3-bromophenyl)-ethyl]-alpha-(2-ethoxyphenyl)-1,2,4-triazole-1-propanenitrileStep 1 - Preparation of 2-(3-bromophenyl)-ethanol. Reduction of 3-bromophenylacetic acid via diborane reduction.

A 2 liter 4 neck flask stirred under nitrogen, equipped with condenser and addition funnel, was charged with 75 gms of 3-bromophenylacetic acid (0.34 mole), in 100 ml of THF. Then 350 ml of 1M diborane-THF complex (0.35 mole) was added dropwise. Upon addition, gas evolved and the reaction was cooled to maintain the temperature below 10°C. After the addition was complete, the reaction was stirred at room temperature until, by thin layer chromatography, the reaction was complete. The reaction was quenched by adding ice water and the product was extracted into ether then washed with 5% NaOH, 5% HCl, water and dried over magnesium sulfate. After concentration, 77.4 gms of product resulted and was chlorinated directly.

NMR (90 MHz): 2.7-2.9(m, 3H), 3.7-3.9(t, 2H) and 7.1-7.4(m, 4H).

Step 2 - Preparation of 2-(3-bromophenyl)ethyl chloride. Chlorination of 2-(3-bromophenyl)ethanol.

A 500 ml 3 neck round bottom flask stirred under nitrogen, equipped with condenser and addition funnel, was charged with 20.0 gms of 2-(3-bromophenyl)ethanol, (0.1 mole, 1.0 eq.) in 60 ml of toluene. Then 22 ml of SOCl_2 (35.9 gms, 0.30 mole, 3.0 eq.) was added dropwise while maintaining the temperature below 15°C with external cooling. Pyridine (8.7 gms, 1.1 eq.) was added dropwise in 10 ml of toluene. The reaction was stirred overnight at room temperature after which thin layer chromatography indicated that the reaction was complete. The mixture was concentrated and the product isolated after adding water, extracting with ether and washing with water. After drying and concentrating, 14.2 gms of an orange oil resulted (64.2% yield).

NMR (60 MHz): 2.8-3.1(m, 2H), 3.5-3.8(m, 2H) and 7.0-7.3(m, 4H).

Step 3 - Preparation of alpha-[2-(3-bromophenyl)ethyl]-2-ethoxyphenylacetonitrile

A 4 neck 1 liter round bottom flask was charged with 3.5 gms of 60% NaH (0.083 mole, 1.4 eq.), washed three times with 25 ml hexanes, in 100 ml of 2:1 toluene:DMF. Then 9.5 gms (0.059 mole, 1.0 eq.) of 2-ethoxybenzyl cyanide in 200 ml of 2:1 toluene:DMF was added and stirred for 2 hours. Then 2-(3-bromophenyl)ethyl chloride (13 gms, 0.059 mole) in 100 ml of 2:1 toluene:DMF was added. The reaction was incomplete after 20 hours and 2.0 gms of 100% KH (0.05 mole) in 25 ml of DMF was added. After four hours, the reaction was complete and then quenched with 10% HCl, extracted with ether, washed with water, dried and concentrated to give 20 gms of a crude oil (97% pure). The product was distilled under reduced pressure and gave 12.3 gms of product having a boiling point of 195-210°C at 1 mm Hg (58.9% yield).

NMR (90 MHz): 1.2-1.4(t, 3H), 2.0-2.3(t, 2H), 2.6-2.9(t, 2H), 3.8-4.2(m, 3H) and 6.7-7.4(m, 8H).

Step 4 - Preparation of alpha-[2-(3-bromophenyl)ethyl]-alpha-(2-ethoxyphenyl)-1,2,4-triazole-1-propanenitrile

A 500 ml 4 neck round bottom flask was charged with 2.8 gms of 100% KH (0.07 mole, 3.0 eq.), washed twice with 25 ml of hexanes, in 50 ml of DMF. To the reaction was added dropwise 8.0 gms of alpha-[2-(3-bromophenyl)ethyl]-2-ethoxyphenyl acetonitrile (0.023 mole, 1.0 eq.) in 100 ml of DMF. After 1 hour, 5.7 gms of chloromethyltriazole-HCl (0.037 mole, 1.6 eq.) was added directly in two portions. After 1 hour, gas liquid chromatography indicated the reaction was incomplete and an additional 2.8 gms (3.0 eq.) of KH was added followed by 1.8 gms (0.5 eq.) of chloromethyltriazole-HCl. After stirring overnight at room temperature, the reaction was complete and quenched by adding a small quantity of MeOH in 100 ml of ether followed by the addition of 10 ml of 10% HCl. Then 200 ml of ether was added then washed four times with 100 ml of water, dried and concentrated to give 7 gms of a crude oil. The product was purified by flash chromatography with 1:1 ethyl acetate:hexane and gave 2.3 gms (24% yield) of a viscous oil.

NMR (60 MHz): 1.4-1.7(t, 3H), 2.1-3.0(m, 4H), 4.0-4.3(m, 2H), 5.0(s, 2H), 6.9-7.4(m, 8H) and 7.8(s, 2H).

Elemental Analysis: $\text{C}_{21} \text{H}_{21} \text{N}_4 \text{BrO}$

Theor: C: 59.28, H: 4.98, N: 13.18, O: 3.76, Br: 18.80
 Found: C: 58.40, H: 5.43, N: 11.56, O: 6.30, Br: 17.69

PREPARATION OF COMPOUND

5 93 - Alpha-[2-(4-methoxyphenyl)ethyl]-alpha-phenyl-1,2,4-triazole-1-propanenitrile

Step 1 - Preparation of alpha-[2-(4-methoxyphenyl)ethyl]-phenylacetonitrile

A 4 neck 2 liter round bottom flask equipped with thermometer, dropping funnel and mechanical stirrer was charged with 117 gms (1.0 mole, 2.0 eq.) of benzyl cyanide and 115 gms (0.5 mole, 1.0 eq.) of 2-(4-methoxyphenyl)ethyl methanesulfonate in 400 ml of DMSO and 200 ml of toluene. The reaction was cooled to 10°C and 50 ms (0.63 mole, 1.2 eq.) of 50% NaOH was added dropwise over 30 minutes. The reaction exothermed to 35°C and was cooled to 10°C. The reaction was stirred for 48 hours at room temperature and gas liquid chromatography indicated 60% product and 40% benzyl cyanide. The reaction was quenched by adding to 2 liters of water, then extracted three times with ether (1 liter), washed twice with water (1 liter) and brine (1 liter). Drying over MgSO₄ and concentrating gave 171 gms of crude oil (67% product). The product was distilled under vacuum and afforded 104 gms of yellow oil (82.5% yield based on mesylate) having a boiling point of 165-170°C at 2 mm Hg.

NMR (60 MHz): 1.9-2.4(m, 2H), 2.5-2.7(m, 2H), 3.7(s, 4H), 6.8-7.2(ABq, 4H) and 7.5(s, 5H).

20 Step 2 - Preparation of 1-bromo-2-cyano-2-phenyl-4-(4-methoxyphenyl)butane

A 4 neck 1 liter round bottom flask was charged with 100 grams (0.40 mole, 1.0 eq.) of alpha-[2-(4-methoxyphenyl)ethyl]-phenylacetonitrile and 139.1 gms of dibromomethane in 200 ml of DMSO. To the reaction was added 79.6 gms of 50% NaOH (0.99 mole, 2.5 eq.) dropwise over 15 minutes. The reaction exothermed to 95°C then was cooled to 50°C and stirred for 1 hour at 50°C. When as liquid chromatography indicated the reaction was complete, the reaction product was poured into 2 liters of water, extracted three times with ether (500 ml), washed with water (1 liter), brine and dried over sulfate. 130 gms (94% yield) of product as an amber oil resulted after concentrating. The product was used directly in the triazole coupling.

NMR (60 MHz): 2.4-2.8(m, 4H), 3.7(s, 2H), 3.8(s, 3H), 6.8-7.3(ABq, 4H) and 7.5-7.8(br s, 5H).

30 Step 3 - Preparation of alpha-[2-(4-methoxyphenyl)ethyl]-alpha-phenyl-1,2,4-triazole-1-propanenitrile

A 1 liter 3 neck flask was charged 57 gms of triazole (0.74 mole, 2.0 eq.) in 300 ml of DMSO and 102 gms of K₂ CO₃ (0.74 mole, 4.0 eq.). The reaction was stirred at 135°C for 1 hour after which 127 gms of 1-bromo-2-cyano-2-phenyl-4-methoxyphenyl)butane (0.37 mole, 1.0 eq.) in 200 ml of DMSO was added. The temperature dropped to 115°C and was heated at 135°C for 2 hours then at 80°C for 18 hours. The reaction was poured into water (3 liters), extracted five times with ethyl acetate (500 ml), washed twice with water (500 ml) and 500 ml of brine. After drying over sulfate, filtering and concentrating gave a yellow oil which was slurried in hexane. A yellow solid formed which was filtered with 10% diethyl ether/90% hexanes. Drying yielded 95 gms of a product having a melting point of 91-94°C (77.4% yield).

NMR (60 MHz): 2.4-2.9(m, 4H), 3.9(s, 3H), 5.1(s, 2H), 6.8-7.4(ABq, 4H), 7.7-7.9(br s, 5H), 8.2(s, 1H) and 8.4(s, 1H).

Elemental analysis: C₂₀ H₂₀ N₄ O

Theor: C: 72.24, H: 6.06, N: 16.87, O: 4.82

Found: C: 72.55, H: 6.39, N: 16.09, O: 5.21

The compounds of the present invention were tested for activity against a number of diseases. The test compounds were dissolved in acetone, methanol and water to form a series of dilutions from 300 ppm to 5 ppm. Depending on when the tests were run, various serial dilutions were used, e.g., 300, 75, 19, 5 or 100, 25, 6. Unless otherwise indicated the plants were sprayed to run-off with a mechanical sprayer the same day or the day before inoculation. The protocol for Wheat Stem Rust (WSR), Wheat Leaf Rust (WLR), Wheat Powdery Mildew (WPM) and Rice Blast (RB) were as follows:

50 WHEAT STEM RUST (WSR)-Puccinia graminis

Wheat seedlings cultivar TYLER were grown in redi-earth and used for screening about seven days after planting. The seedlings were fertilized with LIQUID-M fertilizer prior to use to maintain vigorous plants throughout the testing period.

55 Depending on when the test was run one of three methods was used to prepare the urediospore suspensions:

1. Water/Devilbiss Atomizer: A spore suspension of WSR was prepared by harvesting infected leaves from two to three week old culture plants and shaking the leaves vigorously with water plus TWEEN 80 surfactant. The spore suspension was filtered through cheesecloth to remove debris and adjusted to three to five spores per large square on a hemocytometer. The inoculum was applied to plants by using a Devilbiss atomizer. While the plants were still wet, they were placed in a humidity cabinet.

2. Oil/Devilbiss Atomizer: A spore suspension of WSR and oil was prepared by harvesting spores fresh from infected plants two to three weeks old with a vacuum pump or rust collector or dehydrating frozen spores (from deep freeze) and adding to SOLTROL spray oil at a concentration of five mg spores to one ml oil. The inoculum was applied to the plants with a Devilbiss Atomizer making one pass over the plants

from all four sides. The plants were allowed to dry about twenty minutes and then placed in a humidity cabinet.

3. Oil/Small Special Atomizer: The spore suspension was prepared the same as in method #2 except four mg spores per one ml oil was used. The inoculum was then dispensed into gelatin capsules and applied with a vacuum pump. Four passes were made on both sides of the plant for uniformity. The plants were allowed to dry about twenty minutes and then were placed in a humidity cabinet.

The humidity cabinet supplied 100% free water and was maintained at a constant temperature of 70°F. The inoculated plants were subjected to twelve hours of darkness followed by three to four hours of fluorescent light. The plants were then transferred into the greenhouse and evaluated thirteen days later.

WHEAT LEAF RUST (WLR)-Puccinia recondita

The same procedures used for WSR were used for WLR except that for WLR there was no light requirement.

WHEAT POWDERY- MILDEW (WPM)-Erysiphe graminis

Wheat seedlings cultivar VICTORY 283 were grown in redi-earth. The seedlings were six to seven days old and were fertilized with LIQUID-M fertilizer before use to promote vigorous growth throughout the test period.

The seedlings were inoculated by shaking sporulating culture plants over them, disseminating mildew spores. The inoculated seedlings were placed in subirrigation trays in a controlled temperature room which provided a 70°F environment for disease development.

Since WPM development is greatly affected by volatile chemicals, the pots were spread out as much as possible and the trays are separated according to dose by plastic sheets. Disease development was rated seven to ten days after inoculation on a percent control basis.

RICE BLAST (RB)-Piricularia oryzae

Seedlings of the rice cultivar M-201 were grown in a greenhouse at 20-30°C in 2-inch pots containing unsterilized soil and Turf-Bulder soil/fertilizer for 14 days. The rice plants were not trimmed before use.

Inoculum was produced in-vitro on oatmeal agar (50g Gerber baby oatmeal, 20g bacto agar, 10g bacto dextrose, 10 ml deionized water). The plates were inoculated with a mycelial plug (7-14 days old) of Piricularia oryzae. The outer edge of the dark region was used in the transfer. Inoculated plates were maintained at room temperature under constant fluorescent light.

P. Oryzae plates 10-14 days old were flooded with a solution containing 0.25g sodium oleate, 2g gelatin and 1000ml deionized water. The plates were scraped with a rubber policeman to release conidia, filtered through a double layer of cheesecloth and spore suspension adjusted to 25000-30000 spores/ml using a hemacytometer.

The spore suspension was sprayed on opposite sides of a double row of rice plants using a hand sprayer. Sufficient inoculum was applied to achieve uniform distribution from soil to tip of rice leaves on opposite sides of each pot (approx. 50ml/50 pots). The hand sprayer was shook after each pass to keep the solution in suspension.

The inoculated plants were immediately placed in a humidity cabinet at 25°C for 66 hours prior to moving them to the greenhouse under a plastic tent. The plants were subirrigated but not allowed to stand in water more than 2 hours. The plastic sides of the tent were lifted during work hours and closed at end of day.

After 76 hours under greenhouse conditions the bioassay plants were observed and the percent disease control (as compared to inoculated control) was estimated.

The compounds were tested at different dose rates depending on when the tests were run. The results of the tests are set forth in Table 6 for one dose rate for each compound. If the compound was tested more than once at the dose rate, the average is reported.

TABLE 6
FUNGICIDAL ACTIVITY

% Control at 300 ppm

	<u>WSR</u>	<u>WLR</u>	<u>WPM</u>	<u>RB</u>
1	100	95	99	66
2	97	100	93	83
3	100	100	99	98
4	100	100	100	91
5	92	80*	99	15
6	100	100	96	12
7	100	90*	100	88
8	100	80*	90	95
9	100	50*	97	50
10	100	100**	98	97
11	100	100**	99	100
12	100	90*	90	88
13	94	90*	99	99
14	100	95*	100	100
15	92	50*	70	80
16	100	100**	99	100
17	98	80*	100	93
18	100	95*	100	100
19	99	100**	100	97
20	95	90*	100	85
21	100	90*	100	63
22	100	--	100	50

* At 100 ppm

** At 150 ppm

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% Control at 200 ppm

	<u>WSR</u>	<u>WLR</u>	<u>WPM</u>	<u>RB</u>
23	100	95*	90	0**
24	100	100***	100	40**
25	100	100***	100	50**
26	100	100***	90	83**
27	100	--	90	100
28	100	--	100	0**
29	100	95**	100	90**
30	100	98***	100	60
31	100	95*	100	0
32	100	91***	100	80
33	100	91***	100	0
34	100	80*	--	100
35	100	--	--	100
36	100	80*	100*	87
37	100	91***	100	80
38	100	90*	100	0
39	100	90*	100	0
40	--	100	100	80
41	--	100	100	0
42	99**	95	100	0
43	--	100	100	0
44	100**	99	100	95
45	--	99	100	50
46	--	75	75	50
47	100**	75	95	50
48	--	75	75	0
49	100**	95	99	0
50	100**	75	100	0
51	--	75	100	50
52	--	50	--	90
53	--	95	95	90

0 251 775

% Control at 200 ppm

	<u>WSR</u>	<u>WLR</u>	<u>WPM</u>	<u>RB</u>
54	--	95	95	80
55	--	95	100	80
56	--	99	99	100
57	--	50	50	0

* At 100 ppm

** At 150 ppm

*** At 75 ppm

% Control at 100 ppm

	<u>WSR</u>	<u>WLR</u>	<u>WPM</u>	<u>RB</u>
58	--	0	75	0
59	--	75	95	80
60	--	0	75	0
61	--	0	85	40
62	--	75	95	0
63	--	50	75	0
64	100	99	87	50
65	100	95	97	90
66	--	75	95	90
67	95	99	85	85
68	95	95	99	85
69	100	99	100	87
70	--	50	95	80
71	91	50	99	90
72	97	0	97	92
73	--	0	85	50
74	100	92	100	0
75	100	90	100	0

5

% Control at 100 ppm

	<u>WSR</u>	<u>WLR</u>	<u>WPM</u>	<u>RB</u>	10
76	100	93	100	0	
77	100	50	98	93	
78	99	75	99	100	15
79	95	75	85	95	
80	100	99	99	0	
81	100	75	99	0	20
82	100	0	95	50	
83	--	75	99	98	
84	100	99	99	90	25
85	100	99	97	0	
86	100	90	99	85	
87	100	0	100	100	30
88	--	90	95	0	
89	99	75	99	40	
90	--	95	95	0	35
91	100	85	95	90	
92	--	95	99	0	
93	50	0	99	100	40
94	--	50	90	--	
95	100	97	99	95	
96	--	95	97	--	45
97	99	50****	100****	99****	
98	99	95	99*****	90	50

**** at 200 ppm

***** at 25 ppm

55

60

The compounds were tested at different dose rates depending on when the tests were run. The results of the tests for the better compounds which were tested are set forth in Table 7 for three rates. If the compound

65

was tested at the same dose rate more than once, the average is reported.

TABLE 7
FUNGICIDAL ACTIVITY

<u>Wheat Stem Rust</u>									
Compound/Dose Rate (in ppm)									
		300	200	100	75	33	25	6	5
1	100				100				98
4	100				100				99
6	100				100				100
10	100				100				99
16	100				100				99
22	100				100				95
23			100			100		100	
24			100			100		90	
25			100			100		100	
26			100			100		100	
27			100			100		100	
33			100			100		100	
37			100			100		100	
78				99			100	100	
85				100			100	100	
86				100			100	100	
95				100			100	99	
98				99			99	90	

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Wheat Leaf Rust

Compound/Dose Rate (in ppm)

	150	100	75	38	25	19	6	5
7		90			80		80	
10	100			100				96
11	100			100				94
12		90			95		80	
16	100			100				100
19	100			100				100
21		90			80		80	
23		95			90		90	
24			100			99		97
25			100			99		93
26			100			99		90
28					95 (20)			95
41			95			97		80
49		95			95		75	
69		99			95		85	
80		99			97		95	
98		95			90		90	

Compound 28 was tested at 20 ppm

Wheat Powdery Mildew

Compound/Dose Rate (in ppm)

	300	200	100	50	38	33	25	12	6	5
5										
	1	99			99					98
10	2	100			99					98
	4	100			100					97
	10	98			97					95
15	19	100			100					96
	28		100			100			90	
	29		100			100			95	
20	30		100			100			100	
	31		100			100			95	
	36			100			100		95	
25	41		100			95			95	
	49			100			100		75	
	76			100			99		97	
30	81			99			92		92	
	82			95			97		90	
	83			99			97		97	
35	85			97			95		95	
	86			99			99		96	
	87			100			95		97	
40	88			95			95		95	
	94			90			95		90	
	95			99			94		96	
45	97		100		99			100		
	98						99		95	

Rice Blast

Compound/Dose Rate (in ppm)

	300	200	100	75	50	25	12	6	5
55									
	7	88		85					80
	17	93		99					78
	18	100		100					85
60									
	97		99		90		0		
	98			90		90		75	

The activity of a number of the compounds of the present invention was tested against Peanut Cercospora (PC), Barley Helminthosporium (BH) and Wheat Septoria Nodorum (SNW). The protocol for these tests was as

follows:

PEANUT CERCOSPOORA OR PEANUT EARLY LEAFSPOT (PC)-*Cercospora Arachidicola*

Cercospora arachidicola was cultured on peanut-oatmeal agar (POA) in petri plates for 14 days under fluorescent lights that were 20 cm above the cultures. The petri plates were inoculated with 0.5 ml of a spore suspension made in sterile water containing a few drops of Tween 80. The spore suspension was subsequently spread over the surface of the POA plate by means of a sterile glass rod bent in the form of a hockey stick. Spores were harvested from the plates by adding deionized water containing a small amount of Tween 80 to the POA plates. The agar surface was scraped with a rubber policeman or similar blunt object. The spore suspension was filtered through cheesecloth to remove mycelial and agar fragments and then adjusted to a concentration of $2-4 \times 10^6$ spores per ml.

Fourteen day old TAMNUT 74 peanut plants were inoculated by spraying the leaves with inoculum until a uniform film of inoculum was observed on the plant. The inoculated plants were incubated in a humid environment at 29.5-32.2°C (85-90°F) for 72 hours. The plants were removed from the humid environment, allowed to dry, and placed in a greenhouse. Treatment comparisons were made 10-14 days after inoculation.

BARLEY HELMINTHOSPORIUM or BARLEY SPOT BLOTCH TEST (BH)-*Helminthosporium sativum*

Seven day old HENREY barley seedlings in 5 cm (2") pots were trimmed 24 hours prior to chemical application to provide a uniform plant height and to facilitate uniform inoculation. The plants were sprayed with the test compound via a settling tower (1.1 ml of the desired concentration for each spray). The spray was allowed to settle in the tower for one minute before removing the plants from the tower. After spraying, the plants were allowed to dry for at least 2 hours in a drying hood before inoculation.

The *Helminthosporium sativum* culture used for inoculum was about 3 weeks old, deep black and sporulating. Five ml of deionized water was added to each Petri dish of culture and the spores were scraped off into the water with a rubber policeman. After scraping, the water was filtered through cheesecloth to remove mycelia and agar pieces. One drop of TWEEN 80 surfactant was added to each 100 ml of spore suspension. The spore concentration was adjusted to 25 spores/ml. Inoculum was applied using a hand sprayer. The inoculated plants were put into a greenhouse after sitting in a humidity cabinet at about 21.1°C (70°F) for 24 hours. The plants were allowed to sit in the greenhouse for 6 days prior to scoring the test. The disease was assessed according to "A MANUAL OF ASSESSMENT KEYS FOR PLANT DISEASES" by Clive James (Key No 1.6.1).

WHEAT GLUME BLOTCH (SNW)-*Septoria nodorum*

The inoculum was produced by placing pieces of sporulating mycelium from 3 week old plates on Czapek-Dox V-8 plates or placing a piece of sporulating mycelium in a test tube containing 20 ml of sterile deionized water, shaking well and after 5 minutes placing enough liquid spore mixture in a fresh plate to make a fine film over the plate. The plates were incubated for 48 hours at 20°C in darkness until a white mycelium was observed. Then plates were incubated at 21°C under continuous fluorescent light for 15-20 days. The mycelium had a pink color at the end of the incubation period.

The sporulating plates were flooded with deionized water. The spores were scraped into the water with a rubber policeman. The flooding and scraping was repeated 2-3 times with each plate. The spore suspension was then filtered with cheesecloth. The final spore concentration was adjusted 150-300 spores/ml. Two drops of TWEEN 80 per 500 ml of spore suspension were added.

LEN wheat plants were inoculated by spraying the leaves with the spore suspension and TWEEN 80 using a hand sprayer after optionally spraying the plants with a light mineral oil lightly and waiting 5 minutes. The inoculated plants were incubated for 72 hours in a humidity cabinet at 20°C with a photoperiod of 16 hours light/8 hours dark. The plants were then placed in a growth chamber for 7-9 days at 20°C with a photoperiod of 16 hours light/8 hours dark and then evaluated for percent control.

The compounds were tested at different dose rates depending on when the tests were run. The results of the tests for the better compounds which were tested are set forth in Table 8 for three rates. If the compound was tested at the same dose rate more than once, the average is reported.

TABLE 8
FUNGICIDAL ACTIVITY

Peanut Cercospora

Compound/Dose Rate (in ppm)

	300	75	5
4	100	100	97
6	100	100	100
7	100	100	85
8	98	100	85
11	100	100	100

Barley Helminthosporium

Compound/Dose Rate (in ppm)

	300	75	5
3	100	99	87
10	99	99	84
11	100	99	76
16	100	100	96
19	99	99	100
25	94	94	94
26	94	97	86

Wheat Septoria Nodorum

Compound/Dose Rate (in ppm)

	80	20	5
10	100	100	90
49	100	100	90

COMPARATIVE TESTINGCompounds 2 and 10 versus comparative Compounds C2a, C2b and C10

Side by side in vitro comparative testing of Compounds 2, 10, C2a, C2b and C10 were done against P. herpotrichoides and Septoria tritici. The tests were performed by routine PDA poison agar as follows:

POISON AGAR TEST

Thirty-nine grams of potato dextrose agar (PDA) purchased from Difco was suspended in 1 liter of water. The media was autoclaved at 15 psi for 15 minutes. After autoclaving, the media was allowed to cool for 15 minutes. Then a dilution series was prepared by adding a known amount of fungicide to the molten agar. The fungicide was dissolved in methanol, acetone or DMSO previous to mixing with the agar. The fungi were rated for growth after the following incubation periods:

Pseudocercospora herpotrichoides - 12 days at room temp.

Septoria tritici - 14 days at room temp.

The results were reported in millimeters of radius and the EC_{75} was calculated. The EC_{75} values in ppm were as follows:

Pseudocercospora

C2a	C2b	2	
(phenyl)	(benzyl)	(phenethyl)	
20	12	0.6	
	C10	10	
	(benzyl)	(phenethyl)	
	20	0.2	

Septoria tritici

C2a	C2b	2	
8	10	0.2	
	C10	10	
	4	0.02	

Therefore, the compound of the present invention has an EC_{75} at least 20 times less than the corresponding phenyl or benzyl compounds.

Compound 6 versus comparative Compound C6

In vivo side by side comparative testing of Compound 6 and comparative Compound C6 was also performed.

Control of Barley Spot Blotch

High volume sprays in acetone:methanol:water were applied on an overhead mechanical sprayer to Pennrad barley seedlings in 3 inch pots. The EC_{75} values were calculated and are set forth in Table 9 below. As shown, the compound of the present invention has an EC_{75} more than 20 times less than the corresponding benzyl compound when treated the day of inoculation.

Control of Wheat Leaf Rust with Foliar Sprays

The compounds were suspended in acetone:methanol:water and were sprayed to run-off on an overhead mechanical sprayer onto Pennol wheat seedlings in 3 inch pots. The plants were inoculated with aqueous suspensions of urediospores (20,000/ml) of Puccinia recondita and were incubated for 24 hours at 70°F in mist and for a further 7 days in a greenhouse. The EC_{75} values were calculated and are set forth in Table 9 below.

Control of Wheat Powdery Mildew with Follar Sprays

Routine follar spray using Pennol wheat seedlings in 3 inch pots were performed. The EC₇₅ values in ppm were calculated and are set forth in Table 9 below.

TABLE 9

	<u>Compound 6</u> <u>(phenethyl)</u>	<u>Compound C6</u> <u>(benzyl)</u>
<u>BARLEY SPOT BOTCH</u>		
<u>Initial</u> (inoculated day of treatment)	17	350
<u>WHEAT LEAF RUST</u>		
<u>Curative</u> (inoculated 1 day before treatment)	23	10
<u>Initial</u>	15	5
<u>WHEAT POWDERY MILDEW</u>		
<u>Initial</u>	1	4

As can be seen from the above data, the compound of the present invention clearly controls barley spot blotch better than the comparative compound. Although Compound 6 of the present invention is not one of the better compounds against wheat leaf rust or wheat powdery mildew and the strength of the benzyl series is against the wheat fungi, compound 6 is only somewhat poorer in controlling wheat leaf rust than the comparative benzyl compound and Compound 6 is better against wheat powdery mildew when inoculated on the day of treatment.

Compound 10 versus comparative Compound C10Control of Wheat Powdery Mildew and Wheat Leaf Rust

On two occasions in vivo side by side comparative testing of Compound 10 and comparative Compound C10 was performed using the protocol set forth before Table 6. The results of the two tests are set forth below:

	<u>Test 1</u>	<u>WPM</u>			<u>WLR</u>		
		<u>Compound/Dose rate in ppm ai</u>					
		<u>100</u>	<u>25</u>	<u>6</u>	<u>100</u>	<u>25</u>	<u>6</u>
	10	99	95	95	100	100	99
	C10	95	95	85	95	85	85
<u>Test 2</u>							
	10	100	95	95	100	100	99
	C10	100	95	95	95	85	0

Compound 10 is superior to the comparative compound against wheat leaf rust and as good or better than the comparative compound against wheat powdery mildew.

Based on the foregoing comparative tests the phenethyl compounds are overall superior to corresponding

benzyl and phenyl compounds. The compounds of the present invention are more than one magnitude better against Wheat Foot Rot (*Pseudocercospora*), Wheat Leaf Blotch (*Septoria tritici*) and Barely Spot Blotch (*Helminthosporium sativum*), and generally superior against Wheat Leaf Rust (*Puccinia recondita*) and Wheat Powdery Mildew (*Erysiphe graminis*).

The clear advantage of the phenethyl triazoles of the present invention is their overall superior efficiency against a number of fungi. The preferred compounds of the present invention have good fungicidal activity against Barely Spot Blight, Peanut Early Leafspot and Rice Blast as well as Wheat Leaf Rust and Wheat Stem Rust.

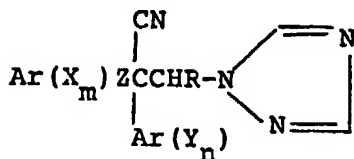
In its mechanical aspects a process of the invention for improving the commercial value and/or profitability of vendible crops from plants whose growth is affected or likely to be affected by fungi comprises (1) charging to a container, fumigation device or mechanical dissemination device a fungicidal composition of the invention as hereinbefore described (2) using the container, fumigator or mechanical dissemination device to apply the fungicidal composition, in the form of granules, dust, smoke, vapour or surfactant-containing liquid preparation to growing plants, to plant seeds or to a growth medium where the plants are growing or are to be grown, or to the fungus itself, (3) controlling the dose of the active ingredient during this application step so that the rate of application of active fungicidal compound is sufficient to combat the fungus but is insufficient to cause an unacceptably adverse effect on the crop plants growing or to be grown in the treated area.

The following words are trademarks which may be registered in some or all of the designated states:

HISIL, Zeolex, Systhane, Liquid-M, Tween, Softrol, Gerber, Turf-Builder.

Claims

1. A fungicidal compound which is a substituted triazole of the formula



wherein

Z is an ethylene group, an ethenylene group, or ethynylene group or an isopropylene group, which ethylene, ethenylene or isopropylene groups may optionally be halogenated;

Ar(X_m) and Ar(Y_n) are substituted or unsubstituted C₆-C₁₀ aromatic ring structures, or Ar(Y_n) may be a five member ring having 4 carbon atoms and one nitrogen, oxygen or sulfur atom, or a six member ring having 1 nitrogen and 5 carbon atoms or 2 nitrogen and 4 carbon atoms;

X and Y are the same or different and may be halogen, (C₁-C₆) alkyl optionally substituted with up to 3 halogens, (C₂-C₆) alkenyl optionally substituted with up to 3 halogens, hydroxy, (C₁-C₆) alkoxy, (C₂-C₆) alkenoxy, phenyl optionally substituted with one or 2 halogens, cyano, amino, mono (C₁-C₆) alkylamino or dialkylamino having independently 1 to 6 carbon atoms in each alkyl group, -C(O)H, SO₂Z wherein Z is (C₁ to C₆) alkyl or aryl and a is 0, 1 or 2, and -C(O)NR₁ R₂ wherein R₁ and R₂ are, independently hydrogen or (C₁-C₆) alkyl;

R is hydrogen or phenyl optionally substituted with up to 3 halogen atoms, a trifluoromethyl group or a (C₁-C₆) alkyl group; and

m and n are independently 0 to 3;

the agronomically acceptable enantiomorphs, acid addition and metal salt complexes thereof.

2. A compound of claim 1 wherein

Z is -CH₂ CH₂ -

Ar(X_m) is an, optionally substituted, phenyl or naphthyl;

Ar(Y_n) is an, optionally substituted, phenyl;

X and Y may be the same or different and are halogen, trifluoromethyl, hydroxy, methoxy, ethoxy, propoxy, methyl, ethyl, phenyl or naphthyl; and

m and n are the same or different and are 0, 1 or 2.

3. A compound of claim 1 wherein Z is -CH=CH- and Ar(X_m), Ar(Y_n), X, Y, m and n are as defined in claim 2.

4. A compound of claim 3 wherein Z is the (E) isomer.

5. The compound of claim 1 or 2 wherein

X_m is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 3-trifluoromethyl, 4-trifluoromethyl, 4-methoxy, 4-methyl, 4-ethyl and 3,4-halogen; and

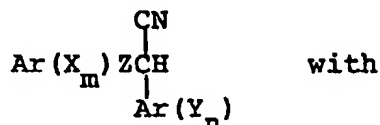
Y_n is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 2,6-halogen, 3-trifluoromethyl, 2-methoxy and 2-ethoxy.

6. A compound of claim 2 wherein (a) X_m is hydrogen and Y_n is 4-halogen or (b) X_m is 4-halogen and Y_n is hydrogen or 4-halogen or 2-halogen or 2-methoxy or 2-ethoxy or (c) X_m is 2-halogen and Y_n is hydrogen or 4-halogen or (d) X_m is 3-halogen or 4-halogen and Y_n is 3-halogen or (e) X_m is 3-halogen and Y_n is hydrogen, 3-halogen, 4-halogen, 2-methoxy, 2-ethoxy or hydrogen or 4-halogen or (f) X_m is 4-trifluoromethyl and Y_n is hydrogen, 4-halogen, 2-methoxy or 2-ethoxy or (g) the substituted triazole is the one named alpha-[2-(4-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-(4-bromophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-(2-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile or alpha-[2-(3-bromophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-phenyl-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile.

7. A fungicidal composition which comprises an agronomically acceptable diluent or carrier and as an active ingredient a fungicidal compound according to any preceding claim.

8. A method for controlling phytopathogenic fungi which comprises of applying to a plant, to plant seed or to a plant habitat, a fungicidally effective amount of a compound as claimed in any of claims 1 to 6 optionally in a composition as claimed in claim 7.

9. A process for preparing a compound of claim 1 which comprises reacting

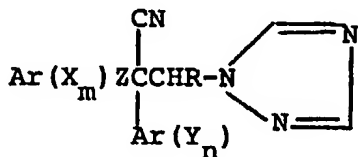


wherein Z is ethylene, ethenylene, ethynylene or isopropylene, Hal means "halogen", X_1 is halogen and Ar, X_m and Y_n are as defined in claim 1.

10. A process for preparing a compound having a quaternary carbon atom which is bonded to an aryl group, a cyano group and a methyltriazole group comprising reacting a compound having a tertiary carbon which is bonded to an aryl group, a cyano group and a hydrogen atom with a halomethyltriazole.

Claims for the following Contracting State: ES

1. The use of a fungicidal compound which is a substituted triazole of the formula



wherein

Z is an ethylene group, an ethenylene group, or ethynylene group or an isopropylene group, which ethylene, ethenylene or isopropylene groups may optionally be halogenated;

$\text{Ar}(X_m)$ and $\text{Ar}(Y_n)$ are substituted or unsubstituted C_6 - C_{10} aromatic ring structures, or $\text{Ar}(Y_n)$ may be a five member ring having 4 carbon atoms and one nitrogen, oxygen or sulfur atom, or a six member ring having 1 nitrogen and 5 carbon atoms or 2 nitrogen and 4 carbon atoms;

X and Y are the same or different and may be halogen, (C_1 - C_6) alkyl optionally substituted with up to 3 halogens, (C_2 - C_8) alkenyl optionally substituted with up to 3 halogens, hydroxy, (C_1 - C_8) alkoxy, (C_2 - C_8) alkenoxy, phenyl optionally substituted with one or 2 halogens, cyano, amino, mono (C_1 - C_8) alkylamino or dialkylamino having independently 1 to 6 carbon atoms in each alkyl group, $-\text{C}(\text{O})\text{H}$, SO_2Z wherein Z is (C_1 to C_8) alkyl or aryl and a is 0, 1 or 2, and $-\text{C}(\text{O})\text{NR}_1\text{R}_2$ wherein R_1 and R_2 are, independently hydrogen or (C_1 - C_8) alkyl;

R is hydrogen or phenyl optionally substituted with up to 3 halogen atoms, a trifluoromethyl group or a (C₁-C₆) alkyl group; and

m and n are independently 0 to 3;

or an agronomically acceptable enantiomorph, acid addition or metal salt complex thereof, together with agronomically acceptable diluent or carrier to make a fungicidal composition.

2. The use according to claim 1 of a compound wherein

Z is -CH₂ CH₂ -

Ar(X_m) is an, optionally substituted, phenyl or naphthyl;

Ar(Y_n) is an, optionally substituted, phenyl;

X and Y may be the same or different and are halogen, trifluoromethyl, hydroxy, methoxy, ethoxy, propxy, methyl, ethyl, phenyl or naphthyl; and

m and n are the same or different and are 0, 1 or 2.

3. The use according to claim 1 of a compound wherein Z is -CH=CH- and Ar(X_m), Ar(Y_n), X, Y, m and n are as defined in claim 2.

4. The use according to claim 1 of a compound as defined in claim 3 wherein Z is the (E) isomer.

5. The use according to claim 1 or 2 of a compound wherein

X_m is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 3-trifluoromethyl, 4-trifluoromethyl, 4-methoxy, 4-methyl, 4-ethyl and 3,4-halogen; and

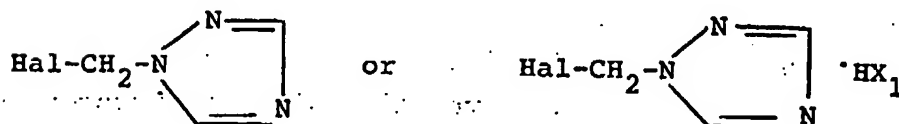
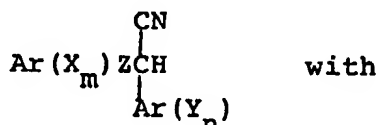
Y_n is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 2,6-halogen, 3-trifluoromethyl, 2-methoxy and 2-ethoxy.

6. The use according to claim 1 of a compound as defined in claim 2 wherein (a) X_m is hydrogen and Y_n is 4-halogen or (b) X_m is 4-halogen and Y_n is hydrogen or 4-halogen or 2-halogen or 2-methoxy or 2-ethoxy or (c) X_m is 2-halogen and Y_n is hydrogen or 4-halogen or (d) X_m is 3-halogen and Y_n is 4-halogen and Y_n is 3-halogen or (e) X_m is 3-halogen and Y_n is hydrogen, 3-halogen, 4-halogen, 2-methoxy, 2-ethoxy or hydrogen or 4-halogen or (f) X_m is 4-trifluoromethyl and Y_n is hydrogen, 4-halogen, 2-methoxy or 2-ethoxy or (g) the substituted triazole is the one named alpha-[2-(4-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-(4-bromophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-(2-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-(2-chlorophenyl)ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile or alpha-[2-(3-bromophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-phenyl-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile.

7. A method for controlling phytopathogenic fungi which comprises of applying to a plant, to plant seed or to a plant habitat, a fungicidally effective amount of a compound as defined in any of claims 1 to 6 optionally in a composition as claimed in any preceding claim.

8. A mechanical process for improving the commercial value and/or profitability of vendible crops from plants whose growth is affected or likely to be affected by fungi which comprises (1) charging to a container, fumigation device or mechanical dissemination device a fungicidal composition using the fungicidal compound according to claim 1 (2) using the container, fumigator or mechanical dissemination device to apply the fungicidal composition, in the form of granules, dust, smoke, vapour or surfactant-containing liquid preparation to growing plants, to plant seeds or to a growth medium where the plants are growing or are to be grown, or to the fungus itself, (3) controlling the dose of the active ingredient during this application step so that the rate of application of active fungicidal compound is sufficient to combat the fungus but is insufficient to cause an unacceptably adverse effect on the crop plants growing or to be grown in the treated area.

9. A process for preparing a compound as described in claim 1 which comprises reacting



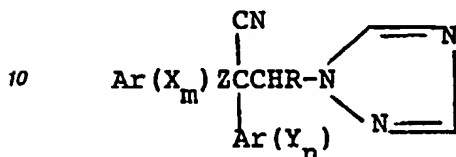
wherein Z is ethylene, ethenylene, ethynylene or isopropylene, Hal means "halogen", X₁ is halogen and Ar, X_m and Y_n are as defined in claim 1.

10. A process for preparing a compound having a quaternary carbon atom which is bonded to an aryl

group, a cyano group and a methyltriazole group comprising reacting a compound having a tertiary carbon which is bonded to an aryl group, a cyano group and a hydrogen atom with a halomethyltriazole.

Claims for the following Contracting State: AT

1. A fungicidal composition which comprises a substituted triazole of the formula



15 wherein

Z is an ethylene group, an ethenylene group, or ethynylene group or an isopropylene group, which ethylene, ethenylene or isopropylene groups may optionally be halogenated;

20 Ar(X_m) and Ar(Y_n) are substituted or unsubstituted C₆-C₁₀ aromatic ring structures, or Ar(Y_n) may be a five member ring having 4 carbon atoms and one nitrogen, oxygen or sulfur atom, or a six member ring having 1 nitrogen and 5 carbon atoms or 2 nitrogen and 4 carbon atoms;

25 X and Y are the same or different and may be halogen, (C₁-C₆) alkyl optionally substituted with up to 3 halogens, (C₂-C₆) alkenyl optionally substituted with up to 3 halogens, hydroxy, (C₁-C₆) alkoxy, (C₂-C₆) alkenoxy, phenyl optionally substituted with one or 2 halogens, cyano, amino, mono (C₁-C₆) alkylamino or dialkylamino having independently 1 to 6 carbon atoms in each alkyl group, -C(O)H, SO₂Ar where Ar is (C₁ to C₆) alkyl or aryl and a is 0, 1 or 2, and -C(O)NR₁ R₂ wherein R₁ and R₂ are, independently hydrogen or (C₁-C₆) alkyl;

R is hydrogen or phenyl optionally substituted with up to 3 halogen atoms, a trifluoromethyl group or a (C₁-C₆) alkyl group; and

30 m and n are independently 0 to 3;

and the agronomically acceptable enantiomorphs, acid addition and metal salt complexes thereof, together with agronomically acceptable diluent or carrier.

2. A composition of claim 1 wherein

Z is -CH₂ CH₂ -

35 Ar(X_m) is an, optionally substituted, phenyl or naphthyl;

Ar(Y_n) is an, optionally substituted, phenyl;

X and Y may be the same or different and are halogen, trifluoromethyl, hydroxy, methoxy, ethoxy, propoxy, methyl, ethyl, phenyl or naphthyl; and

m and n are the same or different and are 0, 1 or 2.

40 3. A composition of claim 1 wherein Z is -CH=CH- and Ar(X_m), Ar(Y_n), X, Y, m and n are as defined in claim 2.

4. A composition of claim 3 wherein Z is the (E) isomer.

5. A composition of claim 1 or 2 wherein

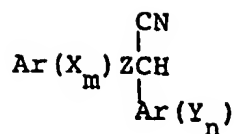
45 X_m is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 3-trifluoromethyl, 4-trifluoromethyl, 4-methoxy, 4-methyl, 4-ethyl and 3,4-halogen; and

Y_n is selected from the group consisting of hydrogen, 2-halogen, 3-halogen, 4-halogen, 2,6-halogen, 3-trifluoromethyl, 2-methoxy and 2-ethoxy.

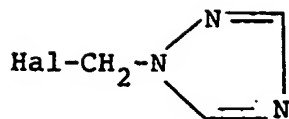
6. A composition of claim 2 wherein (a) X_m is hydrogen and Y_n is 4-halogen or (b) X_m is 4-halogen and Y_n is hydrogen or 4-halogen or 2-halogen or 2-methoxy or 2-ethoxy or (c) X_m is 2-halogen and Y_n is hydrogen or 4-halogen or (d) X_m is 3-halogen or 4-halogen and Y_n is 3-halogen or (e) X_m is 3-halogen and Y_n is hydrogen, 3-halogen, 4-halogen, 2-methoxy, 2-ethoxy or hydrogen or 4-halogen or (f) X_m is 4-trifluoromethyl and Y_n is hydrogen, 4-halogen, 2-methoxy or 2-ethoxy or (g) the substituted triazole is the one named alpha-[2-(4-chlorophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-(4-bromophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-(2-chlorophenyl)-alpha-[2-(4-chlorophenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile or alpha-[2-(4-chlorophenyl)ethyl]-alpha-(2-methoxyphenyl)-1H-1,2,4-triazole-1-propanenitrile or alpha-[2-(3-bromophenyl)ethyl]-alpha-phenyl-1H-1,2,4-triazole-1-propanenitrile or alpha-phenyl-alpha-[2-(4-trifluoromethylphenyl)ethyl]-1H-1,2,4-triazole-1-propanenitrile.

7. A method for controlling phytopathogenic fungi which comprises of applying to a plant, to plant seed or to a plant habitat, a fungicidally effective amount of a compound as defined in any of claims 1 to 6 optionally in a composition as claimed in any preceding claim.

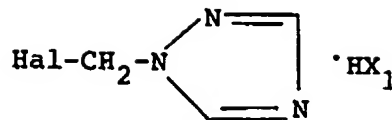
8. A process for preparing a compound as defined in claim 1 which comprises reacting



with



or



wherein Z is ethylene, ethenylene, ethynylene or isopropylene, Hal means "halogen", X_1 is halogen and Ar, X_m and Y_n are as defined in claim 1.

9. A process for preparing a compound having a quaternary carbon atom which is bonded to an aryl group, a cyano group and a methytriazole group comprising reacting a compound having a tertiary carbon which is bonded to an aryl group, a cyano group and a hydrogen atom with a halomethytriazole.